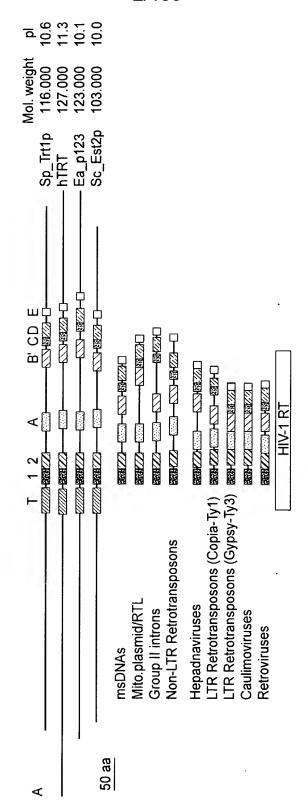
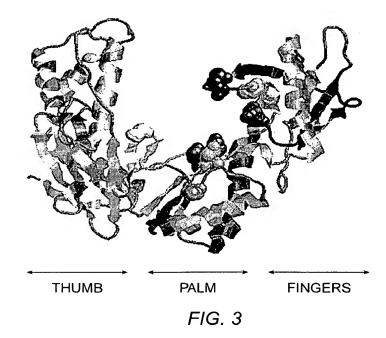


ISEIEWLVLGKRSNAKMCLSDFEKRKQIFAEFIYWLYNSFIIPILQSFFYITESSDLRNR RIIAIPCRGADEEEFTIYKENHKNAIQPTQKILEYLRNKRPTSFTKIYSPTQIADRIKEF KQRLLKKFNNVLPELYFMKFDVKSCYDSIPRMECMRILKD-ALKNENGFFVRSQYFFNTN AKFLHWLMSVYVVELLRSFFYVTETTFOKNR LKDFRWLFISD----IWFTKHNFENLNQLAICFISWLFRQLIPKIIQTFFYCTEISSTVT-TREISWMQVET-SAKHFYYFDHEN-IYVLWKLLRWIFEDLVVSLIRCFFYVTEQQKSYSK LFFYRKSVWSKLQSIGIRQHLKRVQLRDVSEAEVRQHREARPALLTSRLRFIPKP--DGL TVYFRKDIWKLLCRPFI-TSMKMEAFEKINENNVRMDTQK-TTLPPAVIRLLPKK--NTF IVYFRHDTWNKLITPFIVEYFKTYLVENNVCRNHNSYTLS--NFNHSKMRIIPKKSNNEF TYYYRKNIWDVIMKMSI-ADLKKETLAEVQEKEVEEWKKS-LGFAPGKLRLIPKK--TTF RLITN-LRKRFLIKMGSNKKMLVSTNQTLRPVASILKHLINEESSGIPFNLEVYMKLLTF RPIMTFNKKIVNSDRKTTKLTNTKLLNSHLMLKTLKN-RMFKDPFGFAVFNYDDVMKKY KKDLLKHRMFGR-KKYFVRIDIKSCYDRIKQDLMFRIVKK-KLKDPEFVIRKYATIHATS EEFVCKWKQVGQPKLFFATMDIEKCYDSVNREKLSTFLKTTKLLSSDFWIMTAQILKRKN Motif 1 Motif 0 RPIVNMDYVVGARTFRREKRAERLTSRVKALF-SVLMYERA Motif 3 (A) *** Motif 2 * human human human tezl EST2 p123 tez1 EST2 p123 EST2 p123 EST2 p123 tez1 tez1









| Mortal | Immortal | 1 2 3 4 5 6 7 8 9 10 11 12 | hTRT | hTRT | TP1 | GAPDH



Motif 1

O AKKFLNLSLRGFEKHNFSTSLEKTVI 17 KKRMPFFGFSV 181 0 AKIFLRILVRGVPEYGCVVNLRKTVV 19 HGLFPWCGLLL 197 100 89 25 55 YVRYADDILIGYLGSKN 2 KMIKRDLNNFLNS-LGLTMNEEKTLI 4 ETPARFLGYNI 7 LSTYADDTIVLSSDILA 6 NENYLKTFSDWADKWGISVNAAKTGH 25 ESKQSYLGVIL 0 EPPFLWMGITL kfykotkoirógolcussilssfyyatLeesslgfl 14 imrijdDyllítyenn o aulfieklinusrengfkfnmkklot 23 odycdwigisi kcyiredgifyggslsapiudluyddLiefysefk 8 ilkladDfliistdooo o unikklamggfokynakanrdKila 20 keleuwkhsst Ч 1 FRIIAIPCRGADEEFFIIVKENHKNAIQP 42 VLPELYFMKFDVKSCYDSIPRMECMRILKDALKN GQPKLFFATMDIEKCYDSVNREKLSTFLKTTKLLYRPISLLSGLSKMFERLLLKRLFRVDLFK 32 RKEYCSAVFLDISEAFDRVWHEGLLLKLAKILPY WRKLVDFRELNKRTQDFWEVQLGIPHPAG 0 LKKKKSVTVLDVGDAYFSVPLDEDFRKYTAFTIP FGRKKYFVRIDIKSCYDRIKQDLMFRIVKKKLKD PPPELYFVKVDVTGAYDTIPQDRLTEVIASIIKP 0 IRPLSVGNPRDKIVQEVWRMILDTIFDKK 27 FGGSNWFIEVDLKKCFDTISHDLIIKEL \mathbf{K} RYISD hhK Motif E 4 IYQYMDDLYVGSDLEIG 1 HRTKIEELRQHLLRWGLTTPDKKHQK c**k** h CXD I N C ĞΧ Gh h h G c p PcLYFh hDh hDh Motif D ㅁ 40 0 LRPIVNMDYVVGARTFRREKRAERLTSRV 45 41 O FRLITNLRKRFLIKMGSNKKMLVSTNQTL O FRPIMTFNKKIVNSDRKTTKLTTNTKLLN 1 WRKLVDFRELNKRTQDFWEVQLGIPHPAG 6 LLRVVDDFLFITVNKKD 5 LLRLVDDFLLVTPHLTH 10 ω ω WLYNSFIIPILQSFFYITESSDLRNRTVYFRKDIWKLLCRPFITSMKM WLMSVYVVELLRSFFYVTETTFQKNRLFFYRKSVWSKLQSIGIRQHLK WIFEDLVVSLIRCFFYVTEQQKSYSKTYYYRKNIWDVIMKMSIADLKK 366 WLFRQLIPKIIQTFFYCTEISSTVT-IVYFRHDTWNKLITPFIVEYFR LLRL DDFLHIT h Y DDhhh × Motif C Ч TYHKPMLGLPQGSLİSPILCNIVMTLVDNWLEDYI RAGQIGAGVPQGSNLGPILYSIFSSDMPLPHIYHP GIRYQYNVLPQGWKGSPAIFQSSMTKILEPFKKQN SQYLQKVGIPQGSILSSFLCHFYMEDLIDEYLSFT KSYVQCQGIPQGSILSTLLCSLCYGDMENKLFAGI FRhI 9 SILRIGYYPDAWKHAQVKMIL**K**PGKS EGKISKIGPENPYNTPVFAIK**K**KDST CRNHNSYTLSNFNHSKMRIIPKKSNN LSNELGTGKFKFKPMRIVNIPKPKGG h hrhipkk p NNVRMDTQKTTLPPAVIRLLP**K**KNT-EVRQHREARPALLTSRLRFIPKPDG-KEVEEWKKSLGFAPGKLRLIPKKTT-GIPQGS LS hL h Y DL p hh h K pP hh Motif 1 Motif B hPog 441 429 546 Ea_p123 Sc_Est2p 1 Sp_Trt1p hTRT Ea_p123 Sc_Est2p Ea_p123 Sc_Est2p Sp Trtlp Sp_Trt1p Sc al Dm_TART TRT con TRT con RT con RT con $HI\overline{V}-1$ hrrr



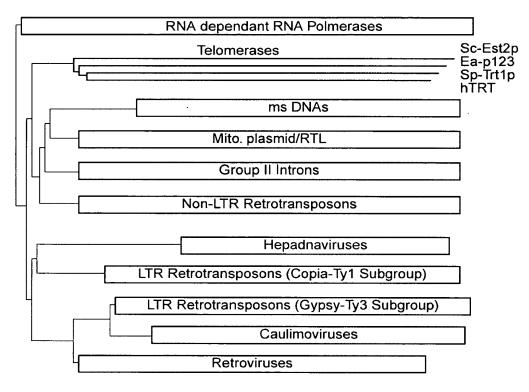


FIG. 6



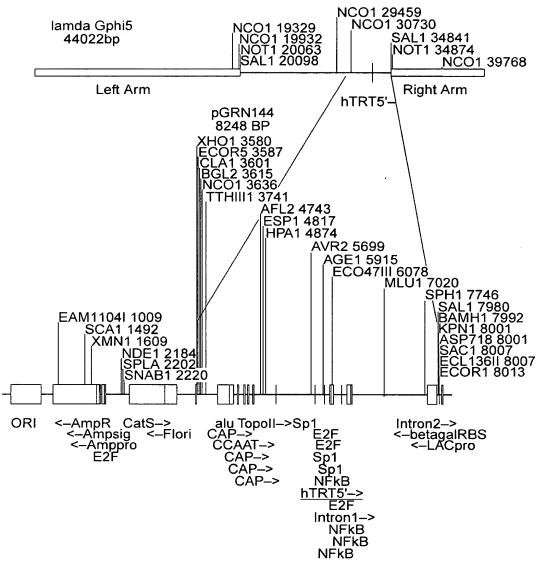


FIG. 7



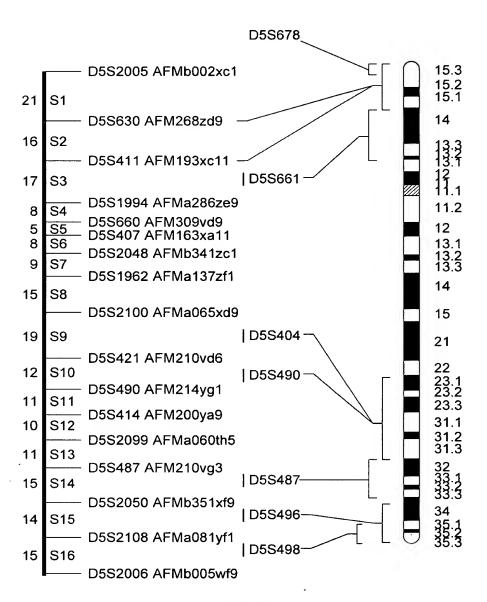
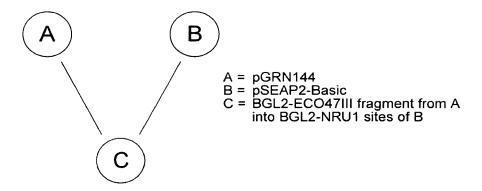


FIG. 8





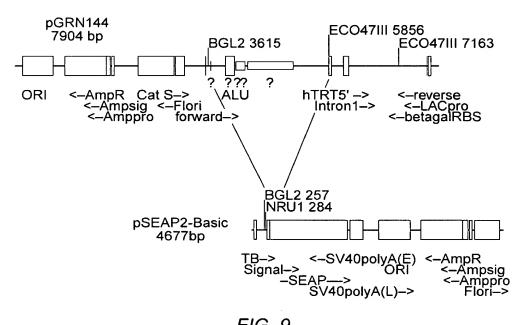


FIG. 9



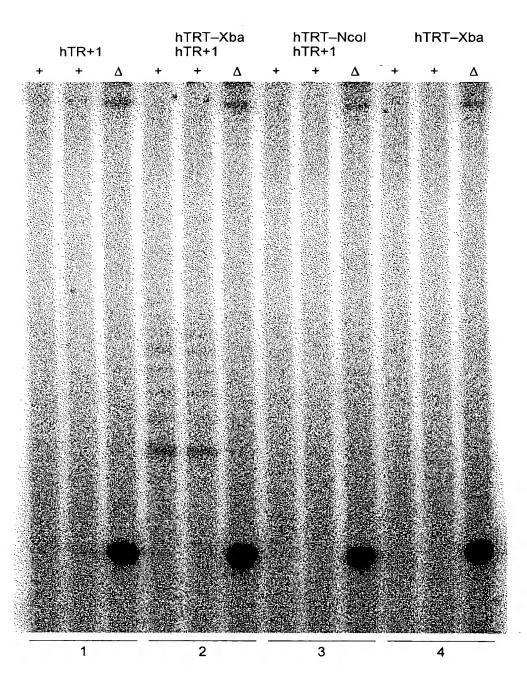


FIG. 10A



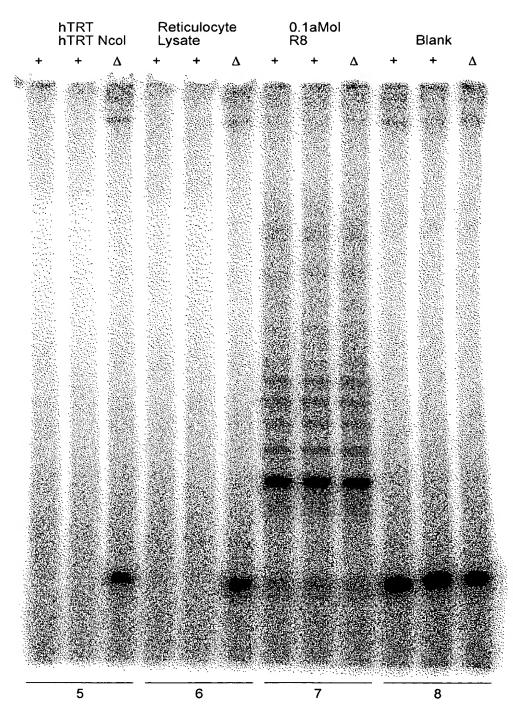


FIG. 10B



Telomerase Specific Motifs

MOTIF I'	E	EAEVR	ENNVR	EKEVE	ENNVC
Σ		13	12	12	σ
	Н	SIGI	RPFI	KMSI	TPFI
	Н	KLO	LLC	VIM	KLI
	Z	SWWS) I WK	IMIN	TWI
	쫐	YRKS	FRKI	YRKI	FRHI
	\succ	LFF	TVY	TYY	IVY
MOTIF T	Wl FFY TE	546 WLMSVYVVELLRSFFYVTETTFOKNRLFFYRKSVWSKLQSIGI 13 EAEVR	429 WLYNSFIIPILQSFFYITESSDLRNRTVYFRKDIWKLLCRPFI 12 ENNVR	441 WIFEDLVVSLIRCFFYVTEQQKSYSKTYYYRKNIWDVIMKMSI 12 EKEVE	366 WLFRQLIPKIIQTFFYCTEISSTVT.IVYFRHDTWNKLITPFI 9 ENNVC
		546	429	441	366
	TRT con	hTRT	\mathtt{spTRT}	Ea p123	Sc_Est2

Telomerase RT Motifs (Fingers)

MOTIF B'	Y q GipQGs 1S l y	69 PELYFVKVDVTGAYDTI 104 YVQCQGIPQGSILSTLLCSLCY	99 YLQKVGIPQGSILSSFLCHFYM	67 PKLFFATMDIEKCYDSV 117 YKQTKGIPQGLCVSSILSSFYY	85 YIREDGLFQGSSLSAPIVDLVY	h hage pp hh
MOTIF A	p lyF D cYD i	69 PELYFVKVDVTGAYDTI	66 RKKYFVRIDIKSCYDRI	67 PKLFFATMDIEKCYDSV 1	68 PELYFMKFDVKSCYDSI	h hDh AF h
MOTIF 2	fR I	0 LRPIV	0 FRLIT	O FRPIM	2 FRIIA	hR h
MOTIF 1	R iPKk	11 SRLRFIPKPDG 0 LRPIV	10	10 GKLRLIPKKTT 0 FRPIM	13 8	p hh h K
	TRT con	hTRT	\mathtt{sptr}	Ea_p123	Sc_Est2	RT con

Telomerase RT Motifs (Palm, Primer Grip)

	100	176	174	141		
MOTIF E	W g s L	FFGFSVNMRSL	WIGISIDMKTL	WKHSSTMINFH	hLG h	
	ć	7 7 7	28	25		
MOT	LILE DDIL IC G n K w g s I	GFEKHNFSTSLEKTVI	LLMRLTDDYLLIT 15 VSRENGFKFNMKKLQT 28 WIGISIDMKTL	GFQKYNAKANRDKILA 25 WKHSSTMNNFH	Gh h cK h	
	г. П	15	15	15		
MOTIF C			LLMRLTDDYLLIT	LILKLADDFLIIS 15	h Y DDhhh	Ĺ
	<u>ر</u>	16	24	18		
E	IKI CON PTPT	SPTRT	Ea_p123	Sc_Est2	RT con	

FIG. 1:



> NFkB_CS1 GGGRQTYYQC NFkB-MHC-I.2 TGGGCTTCCCC

301 GCTGGGGTTGAGGGCGGCCGGGGGGAACCAGCGACATGCGGAGAGCAGCGCAGCGACTCCGACCCCAACTCCCGCCGGCCCCCTTGGTCGCTGTACGCCTCTCGTCGCGTCCGCTGAG

NFkB_CS1
GGGRQTYYQC
NFkB_CS2
RGGGRMTYYCC
Topo_II_cleavage_si

Topo_II_cleavage_site RNYNNCNNGYNGKTNYNY *******

361 AGGGCGCTTCCCCCGCAGGTGTCCTGCCTGAAGGAGCTGGTGGCCCGAGTGCTGCAGAGG TCCCGCGAAGGGGGCGTCCACAGGACGGACTTCCTCGACCACCGGGCTCACGACGTCTCC



1	AAAACCCCAA	AACCCCAAAA	CCCCTTTTAG	AGCCCTGCAG	TTGGAAATAT
51	AACCTCAGTA	TTAATAAGCT	CAGATTTTAA	ATATTAATTA	CAAAACCTAA
101	ATGGAGGTTG	ATGTTGATAA	TCAAGCTGAT	AATCATGGCA	TTCACTCAGC
151	TCTTAAGACT	TGTGAAGAAA	TTAAAGAAGC	TAAAACGTTG	TACTCTTGGA
201	TCCAGAAAGT	TATTAGATGA	AGAAATCAAT	CTCAAAGTCA	TTATAAAGAT
251	TTAGAAGATA	TTAAAATATT	TGCGCAGACA	AATATTGTTG	CTACTCCACG
301	AGACTATAAT	GAAGAAGATT	TTAAAGTTAT	TGCAAGAAAA	GAAGTATTTT
351	CAACTGGACT	AATGATCGAA	CTTATTGACA	AATGCTTAGT	TGAACTTCTT
401	TCATCAAGCG	ATGTTTCAGA	TAGACAAAAA	CTTCAATGAT	TTGGATTTCA
451	ACTTAAGGGA	AATCAATTAG	CAAAGACCCA	TTTATTAACA	GCTCTTTCAA
501	CTCAAAAGCA	GTATTTCTTT	CAAGACGAAT	GGAACCAAGT	TAGAGCAATG
551	ATTGGAAATG	AGCTCTTCCG	ACATCTCTAC	ACTAAATATT	TAATATTCCA
601	GCGAACTTCT	GAAGGAACTC	TTGTTCAATT	TTGCGGGAAT	AACGTTTTTG
651	ATCATTTGAA	AGTCAACGAT	AAGTTTGACA	AAAAGCAAAA	AGGTGGAGCA
701	GCAGACATGA	ATGAACCTCG	ATGTTGATCA	ACCTGCAAAT	ACAATGTCAA
751	GAATGAGAAA	GATCACTTTC	TCAACAACAT	CAACGTGCCG	AATTGGAATA
801	ATATGAAATC	AAGAACCAGA	ATATTTTATT	GCACTCATTT	TAATAGAAAT
851	AACCAATTCT	TCAAAAAGCA	TGAGTTTGTG	AGTAACAAAA	ACAATATTTC
901	AGCGATGGAC	AGAGCTCAGA	CGATATTCAC	GAATATATTC	AGATTTAATA
951	GAATTAGAAA	GAAGCTAAAA	GATAAGGTTA	TCGAAAAAAT	TGCCTACATG
1001	CTTGAGAAAG	TCAAAGATTT	TAACTTCAAC	TACTATTTAA	CAAAATCTTG
1051	TCCTCTTCCA	GAAAATTGGC	GGGAACGGAA	ACAAAAAATC	GAAAACTTGA
1101	TAAATAAAAC	TAGAGAAGAA	AAGTCGAAGT	ACTATGAAGA	GCTGTTTAGC
1151	TACACAACTG	ATAATAAATG	CGTCACACAA	TTTATTAATG	AATTTTTCTA
1201	CAATATACTC	CCCAAAGACT	TTTTGACTGG	AAGAAACCGT	AAGAATTTTC
1251	AAAAGAAAGT	TAAGAAATAT		ACAAGCATGA	ACTCATTCAC
1301	AAAAACTTAT	TGCTTGAGAA	GATCAATACA	AGAGAAATAT	CATGGATGCA
1351	GGTTGAGACC	TCTGCAAAGC	ATTTTTATTA	TTTTGATCAC	GAAAACATCT
1401	ACGTCTTATG	GAAATTGCTC	CGATGGATAT	TCGAGGATCT	CGTCGTCTCG
1451	CTGATTAGAT	GATTTTTCTA	TGTCACCGAG	CAACAGAAAA	GTTACTCCAA
1501	AACCTATTAC	TACAGAAAGA	ATATTTGGGA	CGTCATTATG	AAAATGTCAA
1551	TCGCAGACTT	AAAGAAGGAA	ACGCTTGCTG	AGGTCCAAGA	AAAAGAGGTT
1601	GAAGAATGGA	AAAAGTCGCT	TGGATTTGCA	CCTGGAAAAC	TCAGACTAAT
1651	ACCGAAGAAA	ACTACTTTCC	GTCCAATTAT	GACTTTCAAT	AAGAAGATTG
1701	TAAATTCAGA	CCGGAAGACT	ACAAAATTAA	CTACAAATAC	GAAGTTATTG
1751	AACTCTCACT	TAATGCTTAA	GACATTGAAG	AATAGAATGT	TTAAAGATCC
1801	TTTTGGATTC	GCTGTTTTTA	ACTATGATGA	TGTAATGAAA	AAGTATGAGG
1851	AGTTTGTTTG			AACCAAAACT	CTTCTTTGCA
1901	ACTATGGATA	TCGAAAAGTG	ATATGATAGT	GTAAACAGAG	AAAAACTATC
1951	AACATTCCTA	AAAACTACTA	AATTACTTTC	TTCAGATTTC	TGGATTATGA
2001	CTGCACAAAT	TCTAAAGAGA	AAGAATAACA	TAGTTATCGA	TTCGAAAAAC
2051	TTTAGAAAGA	AAGAAATGAA	AGATTATTTT	AGACAGAAAT	TCCAGAAGAT
2101	TGCACTTGAA	GGAGGACAAT	ATCCAACCTT	ATTCAGTGTT	CTTGAAAATG
2151	AACAAAATGA	CTTAAATGCA	AAGAAAACAT	TAATTGTTGA	
2201	AGAAATTATT	TTAAGAAAGA	TAACTTACTT	CAACCAGTCA	TTAATATTTG
2251	CCAATATAAT	TACATTAACT	TTAATGGGAA	GTTTTATAAA	CAAACAAAAG
2301	GAATTCCTCA	AGGTCTTTGA	GTTTCATCAA	TTTTGTCATC	ATTTTATTAT
2351	GCAACATTAG	AGGAAAGCTC	CTTAGGATTC	CTTAGAGATG	AATCAATGAA

FIG. 13A





2401	CCCTGAAAAT	CCAAATGTTA	ATCTTCTAAT	GAGACTTACA	GATGACTATC
2451	TTTTGATTAC	AACTCAAGAG	AATAATGCAG	TATTGTTTAT	TGAGAAACTT
2501	ATAAACGTAA	GTCGTGAAAA	TGGATTTAAA	TTCAATATGA	AGAAACTACA
2551	GACTAGTTTT	CCATTAAGTC	CAAGCAAATT	TGCAAAATAC	GGAATGGATA
2601	GTGTTGAGGA	GCAAAATATT	GTTCAAGATT	ACTGCGATTG	GATTGGCATC
2651	TCAATTGATA	TGAAAACTCT	TGCTTTAATG	CCAAATATTA	ACTTGAGAAT
2701	AGAAGGAATT	CTGTGTACAC	TCAATCTAAA	CATGCAAACA	AAGAAAGCAT
2751	CAATGTGGCT	CAAGAAGAAA	CTAAAGTCGT	TTTTAATGAA	TAACATTACC
2801	CATTATTTTA	GAAAGACGAT	TACAACCGAA	GACTTTGCGA	ATAAAACTCT
2851	CAACAAGTTA	TTTATATCAG	GCGGTTACAA	ATACATGCAA	TGAGCCAAAG
2901	AATACAAGGA	CCACTTTAAG	AAGAACTTAG	CTATGAGCAG	TATGATCGAC
2951	TTAGAGGTAT	CTAAAATTAT	ATACTCTGTA	ACCAGAGCAT	TCTTTAAATA
3001	CCTTGTGTGC	AATATTAAGG	ATACAATTTT	TGGAGAGGAG	CATTATCCAG
3051	ACTTTTTCCT	TAGCACACTG	AAGCACTTTA	TTGAAATATT	CAGCACAAAA
3101	AAGTACATTT	TCAACAGAGT	TTGCATGATC	CTCAAGGCAA	AAGAAGCAAA
3151	GCTAAAAAGT	GACCAATGTC	AATCTCTAAT	TCAATATGAT	GCATAGTCGA
3201	CTATTCTAAC	TTATTTTGGA	AAGTTAATTT	TCAATTTTTG	TCTTATATAC
2251	TCCCCTTTTC	CCCTTTTCCC	CHUTTUCCCC		

FIG. 13B

1	MEVDVDNQAD	NHGIHSALKT	CEEIKEAKTL	YSWIQKVIRC	RNQSQSHYKD
51	LEDIKIFAQT	NIVATPRDYN	EEDFKVIARK	EVFSTGLMIE	LIDKCLVELL
101	SSSDVSDRQK	LQCFGFQLKG	NQLAKTHLLT	ALSTQKQYFF	QDEWNQVRAM
151	IGNELFRHLY	TKYLIFQRTS	EGTLVQFCGN	NVFDHLKVND	KFDKKQKGGA
201	ADMNEPRCCS	TCKYNVKNEK	DHFLNNINVP	NWNNMKSRTR	IFYCTHFNRN
251	NQFFKKHEFV	SNKNNISAMD	RAQTIFTNIF	RFNRIRKKLK	DKVIEKIAYM
301	LEKVKDFNFN	YYLTKSCPLP	ENWRERKQKI	ENLINKTREE	KSKYYEELFS
351	YTTDNKCVTQ	FINEFFYNIL	PKDFLTGRNR	KNFQKKVKKY	VELNKHELIH
401	KNLLLEKINT	REISWMQVET	SAKHFYYFDH	ENIYVLWKLL	RWIFEDLVVS
451	LIRCFFYVTE	QQKSYSKTYY	YRKNIWDVIM	KMSIADLKKE	TLAEVQEKEV
501	EEWKKSLGFA	PGKLRLIPKK	TTFRPIMTFN	KKIVNSDRKT	TKLTTNTKLL
551	NSHLMLKTLK	NRMFKDPFGF	AVFNYDDVMK	KYEEFVCKWK	QVGQPKLFFA
601	TMDIEKCYDS	VNREKLSTFL	KTTKLLSSDF	WIMTAQILKR	KNNIVIDSKN
651	FRKKEMKDYF	RQKFQKIALE	GGQYPTLFSV	LENEQNDLNA	KKTLIVEAKQ
701	$\mathtt{RNYFKKDNLL}$	QPVINICQYN	YINFNGKFYK	QTKGIPQGLC	VSSILSSFYY
751	ATLEESSLGF	LRDESMNPEN	PNVNLLMRLT	DDYLLITTQE	NNAVLFIEKL
801	INVSRENGFK	FNMKKLQTSF	PLSPSKFAKY	GMDSVEEQNI	VQDYCDWIGI
851	SIDMKTLALM	PNINLRIEGI	LCTLNLNMQT	KKASMWLKKK	LKSFLMNNIT
901	HYFRKTITTE	DFANKTLNKL	FISGGYKYMQ	CAKEYKDHFK	KNLAMSSMID
951	LEVSKIIYSV	TRAFFKYLVC	NIKDTIFGEE	HYPDFFLSTL	KHFIEIFSTK
1001	KYIFNRVCMI	LKAKEAKLKS	DOCOSLIOYD	A	

FIG. 14



1272 86 1018 1078 1405 1332 901 880 958 actcaataacaataccaagtcaaattccaatatgaaggtgttattagtgatcgataatatttctattttatcggtcgtta ggtaccgatttactttcctttcttcataagctaattgcttcctcgaacgctcctaaatctctggaaatattttacaaga agctcttggagtagctcacagaaatccttacaaatcttctgatgagactatattagattcattacagtccgtgcatattc ccaaatatgtatcatctcgtattaggctttttccgttttactcctggaatcgtacctttttcactattccccctaatga ataatctaaattagtttcgcttataattgatagtagtagaaagattggtgattctactcgtgtaatgttattagtttaaa ccaagtataaggacaaaaagaacaacttccttcccctaaaagacttttactttattaatttacttttcaaatatatttcg ggttcgcttacttttaatcgtggtactgttttagctgctacttctagccaaccgcgtgtttctaccccgtcattggatat ttaacatggagccttacactttagatgagtcacgtcgcatgatggagtatttggtatcatccaacgtttgccttgaaaag gttgataattatttgcaaaatcatgtccttagtggtggtgatccgcgcgaaagttttttgatgcttgcacacgtctagcatg attgagatattcaaaaatttctatccactacaactcctttaacgcgggttttatttttctatttctattctattctcatgttgt gtatatatatttttgttttgatttttttttctattcgggatagctaatatggggcag gtaaggtattctaattgtgaaatatttacctgcaattactgttccaaagaga GTA V TCGAAT AGC AAT N GCA AAT N CGT GAG E 9999 AGA TTT F CGC M ATG AGT S GAA E GAT D GAT CCA AGG. R CAA AAG AGC AAA AGC K S GTA GAT AGA AGT CAG ATG TAT GAC GAA ညည GAT ATG CAT H AAA TCC ACC AAT GAG E GAA ე ეტტ ttgtatttaaccgataaag AAT CAT H TTA CAC H ACC 999 AAT TGTAAA AAT ATG ACC AGC 1019 TAC 21 Y CTGCAT H TAT AAA K 1139 1199 1406 1273 87 1079 1333 401 61 801 81 481 561

FIG. 15A



1721 195 1907 245 1967 265 2027 285 2087 305 2147 325 2207 345 0 0 0 CAA O GTG V AAA K $_{\rm Y}^{\rm TAT}$ AGG R ACA T CAC AAA GTG ATT CCA CTG GTA H K V I P L V TTT AAC F N TGG W CCA P AAT N TTT F gtaaataccggttaagatgttgcgcactttgaacaagactgacaagtatag AT CTA TAT L Y TGG ATT W I GAG E GTG V AAT N gtaactaatactgttatccttcataactaattttag CTT AAT N CTT L AAC N AAG K ATA I AAT N $\mathop{\mathrm{TTG}}_{\mathrm{L}}$ ACT T AGC S TGG W TAC Y AAA K GTG AAG CAA V K Q TAC CTT ATA TCT Y L I S ATT I AAG K AGT S TTT F ATG M GTT V TTT F TCC S AGG R CAC H CTA L CCA CTT P L ACA T AGT S CTC L AGT S CAA O GAA E ATT I CGT R ACA T GGC ATA (AAT N CCC AAA P K AAG CGA ACC ATT K R T I TCA S AAC GCA TTT N A F AAC N CCT P ATT I CGG R AGC . TCT S GAT D O D ACT ATA I AAT N CAA O TGT GAT CAG Q AGA R AAG K TGG W $_{\rm L}^{\rm CTT}$ AAA K TCC AGT S TAC Y AAA K TTT F AAG K AAT N TCA S AAG K CAA O GAA E TCA S 1662 176 1782 216 1842 236 1908 246 1722 196 2028 286

FIG. 15B



2775 495 2835 515 ACT TTC TTA ATA AAG gtattaatttttggtcatcaatgtactttacttctaatctatta F L I K CAG AAA ACT Q K T GAG E GCG A TAT Y GTT ATT CGT CTA TTA CCT AAG AAG AAT ACC TTT CGT CTC ATT ACG V I R L L P K K N T F R L I T T AAC N TTT F CTCAAC N G gtattgtataaaatttattaccactaacgattttaccag AC D ACT T CAA AAA K GAT D AAG K CAA ACT Q T GAA E ATG M CGC R $_{
m L}^{
m TTA}$ AGA R TTT F ATT I AGG R AAA K AAC N ATT I GCG A GGT G GTT V GAG E ACG CCT P GAA E AGT S gtattttaaagtattttttgcaaaagctaatattttcag AAC AAT $_{
m N}$ TTT F ATA I ATG M GTT V AGT S TTA GTC AGT L V S ATG AAA M GAT D ATT ATA I ACT GAA E AGT S TTA AGT L S GAA E AAG K TTT F gtaatatgccaaatttttttaccattaattaacaatcag CGA R TCA S TAC AAT TCG Y N S AAT N AAT N AAC AAA AAA ATG N K K M ATG TGC : CGA R TTT ATT ACA F I T GAT TTA D L A AA TTA AAA L K AGA R CAT H CCA GCA (P GCG A AGT S CCC P 2836 AAT TTA AGA AAA AGA 516 N L R K R ATA I AAT N TAC CGA R TCA S ATA CCT TCA S GAA E TGC ATG M 2776 ACT TTG 496 T L 2466 AAA AGG 406 K R ACT T 2907 ttagcag 525 2586 ATC 446 I GAA E CTC L 2526 426 2646 466

FIG. 15C



3465 679 3532 692 3593 708 3653 728 3777 764 3276 ACA AAA AAC TTT GTT AGT GAG GCG TTT TCC TAT T gtaagtttatttttcattggaattttttaacaa 632 T $\,$ K $\,$ N $\,$ F $\,$ V $\,$ S $\,$ E $\,$ A $\,$ F $\,$ S $\,$ Y $\,$ F GTT AAG gtataccaattgttgaattgtaataaca \ensuremath{V} 3714 GTT AAT AAA AAG GAT GCA AAA AAA TTT TTG AAT TTA TCT TTA AGA G gtgagttgctgtcattcc 749 V N K K D A K K F L N L S L R G GAT ATG GTG CCT TTT GAA AAA GTC GTG CAG TTA CTT TCT ATG AAA ACA D M V P F E K V V Q L L S M K T TTT F 3594 ATT CTG TCA TCT TTT TTG TGT CAT TTC TAT ATG GAA GAT TTG ATT GAA TAC CTA TCG 709 I L S S F L C H F Y M E D L I D E Y L S 3654 TTT ACG AAA AAG AAA GGA TCA GTG TTG TTA CGA GTA GTC GAC GAT TTC CTC TTT ATA ACA 729 F T K K G S V L L R V V D D F L F I T TTT AAT GAA AGC AAG AAA F N E S K K tatataatgcgcgattcctcattattaattttgcag G CGT AAG AAG TAT TTT GTA CGG ATA GAT ATA R K Y F V R I D I GİT GAT TTT GTG GAT TAT TGG ACC AAA AGT TCT TCT GAA ATT V D Y W T K S S S E I ACG AGC CTG GAG AAA ACA T S L E K T 3533 ctaatgaaactag ATA GGA AAT TCT CAA TAC CTT CAA AAA GTT GGT ATC CCT CAG GGC 693 I G N S Q Y L Q K V G I P Q G $\,$ AGT AAT GGG ATA ATA AAC AAT ACT TTT S N G I I N N T F CAT CTC TCT GGA CAC ATT H L S G H I GAG taagttctaaccgttgaag GA AAT AAA ATG CTC AAG GAA K M L K E GAT D TTG TAT Y attcttttttag TT ACT ATA AAC I N TCA S 3406 660

FIG. 15L



4209 888 4274 903 4339 917 4528 966 gcc A TG gtacgtgtc W TTT ATT CCC CAA AGA ATG TTC ATA ACG G gtgagtacttatttaactaga F I P Q R M F I T D GAT TAA tgtcattttcaatttatatatacatcctttattactggtgtcttaaacaatattattactaagtata $_{
m D}$ * TGC TGC AAT ATA TAT AGG CTA GGA TAC TCT ATG TGT ATG AGA GCA CAA GCA TAC TTA AAA C C N I Y R L G Y S M C M R A Q A Y L K AAA K TTT CAG TCA TTG ACT F Q S L T TGG AAA AAG TTG W K K L GCA A ATG M CAT H AAA K TTG AAA K ACA T ACG T AGA R GGA AGA AAA ATT G R K I GAT D CTG L AAT N GAA E ATG M TTC GAA CAG CTA ATA TAC CAA F E Q L I Y Q GAG E GGA G GCA A GTA V CTT ATC AAG CCG CTA AGA CCA GTT TTG CGA CAG GTG L I K P L R P V L R Q V TCT S GAC ATT ACC D I T TCT CTT L AGG R TCT S TCC S TGT C TTG AAT GTT ATT L N V I ACA T $_{
m L}^{
m TTG}$ TTT F TCT S ATT I TTC F $\tt ggtctcgagacttcagcaatattgacacatcag \ G \ CTT \\ L$ AAC N CGT R AG R AGG R 4469 CCC TCT TTC AAA TAT CAT CCA TGC 947 P S F K Y H P C CTT AGT S aaagtcattaattaaccttag AT ACG T GAA E AGG ATG AAG GAT ATA R M K D I TAT Y GAT D GGA G TTA 4210 889

FIG. 15E





1 gcagegetge gteetgetge gcaegtggga agecetggee ceggeeacce eegegatgee 61 gegegeteee egetgeegag eegtgegete eetgetgege ageeactace gegaggtget 121 gccgctggcc acgttcgtgc ggcgcctggg gccccagggc tggcggctgg tgcagcgg 181 ggacccggcg getttccgcg cgctggtggc ccagtgcctg gtgtgcgtgc cctgggacgc 241 acggccgccc cccgccgccc cctccttccg ccaggtgtcc tgcctgaagg agctggtggc 301 ccgagtgctg cagaggctgt gcgagcgcgg cgcgaagaac gtgctggcct tcggcttcgc gctgctggac ggggcccgcg ggggcccccc cgaggccttc accaccagcg tgcgcagcta 421 Cctgcccaac acggtgaccg acgcactgcg ggggagcggg gcgtgggggc tgctgctgcg 481 ccgcgtgggc gacgacgtgc tggttcacct gctggcacgc tgcgcgctct ttgtgctggt 541 ggctcccagc tgcgcctacc aggtgtgcgg gccgccgctg taccagctcg gcgctgccac 601 tcaggcccgg ccccgccac acgctagtgg accccgaagg cgtctgggat gcgaacgggc 661 ctggaaccat agcgtcaggg aggccggggt cccctgggc ctgccagcc cgggtgcgag gaggegeggg ggeagtgeea geegaagtet geegttgeee aagaggeeea ggegtggege 781 tgcccctgag ccggagcgga cgcccgttgg gcaggggtcc tgggcccacc cgggcaggac 841 gcgtggaccg agtgaccgtg gtttctgtgt ggtgtcacct gccagacccg ccgaagaagc 901 cacciette gaggetece tetetegeac gegecactee cacceateeg teggececa 961 geaccaegeg ggeececcat ceacategeg gecaccaegt cectgggaca egeettigtee 1021 cccggtgtac gccgagacca agcacttcct ctactcctca ggcgacaagg agcagctgcg 1081 gccctccttc ctactcagct ctctgaggcc cagcctgact ggcgctcgga ggctcgtgga 1141 gaccatcttt ctgggttcca ggccctggat gccagggact ccccgcaggt tgccccgcct 1201 gececagege taetggeaaa tgeggeeeet gtttetggag etgettggga accaegegea gtgcccctac ggggtgctcc tcaagacgca ctgcccgctg cgagctgcgg tcaccccagc 1321 agccggtgtc tgtgcccggg agaagcccca gggctctgtg gcggcccccg aggaggagga 1381 cacagacccc cgtcgcctgg tgcagctgct ccgccagcac agcagcccct ggcaggtgta 1441 cggcttcgtg cgggcctgcc tgcgccggct ggtgccccca ggcctctggg gctccaggca 1501 caacgaacgo ogottootoa ggaacaccaa gaagttoato toootgggga agcatgocaa 1561 gctctcgctg caggagctga cgtggaagat gagcgtgcgg gactgcgctt ggctgcgcag 1621 gagcccaggg gttggctgtg ttccggccgc agagcaccgt ctgcgtgagg agatcctggc caagtteetg caetggetga tgagtgtgta egtegtegag etgeteaggt etttettta tgteaeggag accaegttte aaaagaaeag getetttte taeeggaaga gtgtetggag 1801 caagttgcaa agcattggaa tcagacagca cttgaagagg gtgcagctgc gggagctgtc
1861 ggaagcagag gtcaggcagc atcgggaagc caggcccgcc ctgctgacgt ccagactccg
1921 cttcatcccc aagcctgacg ggctgcggcc gattgtgaac atggactacg tcgtgggagc 1981 cagaacgttc cgcagagaaa agagggccga gcgtctcacc tcgagggtga aggcactgtt 2041 cagcgtgctc aactacgagc gggcgcggcg ccccggcctc ctgggcgcct ctgtgctggg 2101 cctggacgat atccacaggg cctggcgcac cttcgtgctg cgtgtgcggg cccaggaccc 2161 gccgcctgag ctgtactttg tcaaggtgga tgtgacgggc gcgtacgaca ccatccccca ggacaggete acggaggtea tegecageat cateaaacee cagaacaegt actgegtgeg 2281 toggtatgoc gtggtocaga aggoogocoa tgggoacgto ogoaaggoot toaagagooa 2341 cgtetetace ttgacagace tecageegta catgegacag ttegtggete acctgeagga 2401 gaccageceg etgagggatg cegtegteat egageagage teeteeetga atgaggeeag 2461 cagtggcctc ttcgacgtct tcctacgctt catgtgccac cacgccgtgc gcatcagggg 2761 gacagtggtg aacttccctg tagaagacga ggccctgggt ggcacggctt ttgttcagat 2821 gccggcccac ggcctattcc cctggtgcgg cctgctgctg gatacccgga ccctggaggt 2881 gcagagcgac tactccagct atgcccggac ctccatcaga gccagtctca ccttcaaccg 2941 cggcttcaag gctgggagga acatgcgtcg caaactcttt gggggtcttgc ggctgaagtg 3001 tcacagcctg tttctggatt tgcaggtgaa cagcctccag acggtgtgca ccaacatcta 3061 caagatcctc ctgctgcagg cgtacaggtt tcacgcatgt gtgctgcagc tcccatttca 3121 tcagcaagtt tggaagaacc ccacattttt cctgcgcgtc atctctgaca cggcctccct 3181 ctgctactcc atcctgaaag ccaagaacgc agggatgtcg ctgggggcca agggcgccgc3241 cggccctctg ccctccgagg ccgtgcagtg gctgtgccac caagcattcc tgctcaagct3301 gactcgacac cgtgtcacct acgtgccact cctggggtca ctcaggacag cccagacgca 3361 gctgagtcgg aagctcccgg ggacgacgct gactgccctg gaggccgcag ccaacccggc 3421 actgecetea gaetteaaga eeateetgga etgatggeea eeegeeeaca geeaggeega 3481 gagcagacac cagcagcet gtcacgccgg gctctacgtc ccagggaggg aggggcgcc 3541 cacacccagg cccgcaccgc tgggagtctg aggcctgagt gagtgtttgg ccgaggcctg 3601 catgtccggc tgaaggctga gtgtccggct gaggcctgag cgagtgtcca gccaagggct 3661 gagtgtccag cacacctgcc gtcttcactt ccccacaggc tggcgctcgg ctccacccca 3721 gggccagett tteeteacea ggageeegge tteeacteee cacataggaa tagteeatee 3781 ccagattege cattetteac ecetegeect geecteettt geetteeace eceaceatee 3841 aggtggagae ectgagaagg acceteggag etetgggaat ttggagtgae caaaggtgtg 3901 ccctgtacac aggcgaggac cctgcacctg gatgggggtc cctgtggggtc aaattggggg 3961 gaggtgctgt gggagtaaaa tactgaatat atgagttttt cagttttgaa aaaaa





MPRAPRCRAVRSLLRSHYREVLPLATFVRRLGPOGWRLVORGDP AAFRALVAQCLVCVPWDARPPPAAPSFRQVSCLKELVARVLQRL CERGAKNVLAFGFALLDGARGGPPEAFTTSVRSYLPNTVTDALR GSGAWGLLLRRVGDDVLVHLLARCALFVLVAPSCAYQVCGPPLY QLGAATQARPPPHASGPRRRLGCERAWNHSVREAGVPLGLPAPG ARRRGGSASRSLPLPKRPRRGAAPEPERTPVGQGSWAHPGRTRG PSDRGFCVVSPARPAEEATSLEGALSGTRHSHPSVGRQHHAGPP STSRPPRPWDTPCPPVYAETKHFLYSSGDKEQLRPSFLLSSLRP SLTGARRLVETIFLGSRPWMPGTPRRLPRLPQRYWQMRPLFLEL LGNHAQCPYGVLLKTHCPLRAAVTPAAGVCAREKPOGSVAAPEE EDTDPRRLVQLLRQHSSPWQVYGFVRACLRRLVPPGLWGSRHNE RRFLRNTKKFISLGKHAKLSLQELTWKMSVRDCAWLRRSPGVGC VPAAEHRLREEILAKFLHWLMSVYVVELLRSFFYVTETTFOKNR LFFYRKSVWSKLQSIGIRQHLKRVQLRELSEAEVRQHREARPAL LTSRLRFIPKPDGLRPIVNMDYVVGARTFRREKRAERLTSRVKA LFSVLNYERARRPGLLGASVLGLDDIHRAWRTFVLRVRAQDPPP ELYFVKVDVTGAYDTIPQDRLTEVIASIIKPQNTYCVRRYAVVQ KAAHGHVRKAFKSHVSTLTDLQPYMRQFVAHLQETSPLRDAVVI EQSSSLNEASSGLFDVFLRFMCHHAVRIRGKSYVQCQGIPQGSI LSTLLCSLCYGDMENKLFAGIRRDGLLLRLVDDFLLVTPHLTHA KTFLRTLVRGVPEYGCVVNLRKTVVNFPVEDEALGGTAFVQMPA HGLFPWCGLLLDTRTLEVQSDYSSYARTSIRASLTFNRGFKAGR NMRRKLFGVLRLKCHSLFLDLQVNSLQTVCTNIYKILLLQAYRF HACVLQLPFHQQVWKNPTFFLRVISDTASLCYSILKAKNAGMSL GAKGAAGPLPSEAVQWLCHQAFLLKLTRHRVTYVPLLGSLRTAO TQLSRKLPGTTLTALEAAANPALPSDFKTILD

FIG. 17

 ${\tt TTATGTCACGGAGACCACGTTTCAAAAGAACAGGCTCTTTTTCTACCGGAAGAGTGTCTG}$ GAGCAAGTTGCAAAGCATTGGAATCAGACAGCACTTGAAGAGGGTGCAGCTGCGGGAGCT CCGCTTCATCCCCAAGCCTGACGGGCTGCGGCCGATTGTGAACATGGACTACGTCGTGGG ${\tt AGCCAGAACGTTCCGCAGAGAAAAGAGGGCCGAGCGTCTCACCTCGAGGGTGAAGGCACT}$ GTTCAGCGTGCTCAACTACGAGCGGCGCGCGCCCCGGCCTCCTGGGCGCCTCTGTGCT GGGCCTGGACGATATCCACAGGGCCTGGCGCACCTTCGTGCTGCGTGTGCGGGCCCAGGA CCCGCCGCCTGAGCTGTACTTTGTCAAGGTGGATGTGACGGCGCGTACGACACCATCCC CCAGGACAGGCTCACGGAGGTCATCGCCAGCATCATCAAACCCCAGAACACGTACTGCGT CCACGTCCTACGTCCAGTGCCAGGGGATCCCGCAGGGCTCCATCCTCTCCACGCTGCTCT GCAGCCTGTGCTACGGCGACATGGAGAACAAGCTGTTTGCGGGGGATTCGGCGGGACGGGC TGCTCCTGCGTTTGGTGATGATTTCTTGTTGGTGACACCTCACCTCACCCACGCGAAAA CCTTCCTCAGGACCCTGGTCCGAGGTGTCCCTGAGTATGGCTGCGTGAACTTGCGGA ${\tt AGACAGTGGTGAACTTCCCTGTA\^GAAGACGAGGCCCTGGGTGGCACGGCTTTTGTTCAGA}$ TGCCGGCCCACGGCCTATTCCCCTGGTGCGGCCTGCTGCATACCCGGACCCTGGAGG TGCAGAGCGACTACTCCAGCTATGCCCGGACCTCCATCAGAGCCAGTCTCACCTTCAACC GCGGCTTCAAGGCTGGGAGGAACATGCGTCGCAAACTCTTTGGGGTCTTGCGGCTGAAGT GTCACAGCCTGTTTCTGGATTTGCAGGTGAACAGCCTCCAGACGGTGTGCACCAACATCT ACAAGATCCTCCTGCTGCAGGCGTACAGGTTTCACGCATGTGTGCTGCAGCTCCCATTTC ATCAGCAAGTTTGGAAGAACCCCACATTTTTCCTGCGCGTCATCTCTGACACGGCCTCCC TCTGCTACTCCATCCTGAAAGCCAAGAACGCAGGGATGTCGCTGGGGGCCCAAGGGCCCCG CCGGCC7TCTGCCCTCCGAGGCCGTGCAGTGGCTGTGCCACCAAGCATTCCTGCTCAAGC TGACTCGACACCGTGTCACCTACGTGCCACTCCTGGGGTCACTCAGGACAGCCCAGACGC AGCTGAGTCGGAAGCTCCCGGGGACGACGCTGACTGCCCTGGAGGCCGCAGCCAACCCGG CACTGCCCTCAGACTTCAAGACCATCCTGGACTGATGGCCACCCGCCCACAGCCAGGCCG CCACACCCAGGCCTGCACCGCTGGGAGTCTGAGGCCTGAGTGATTTTGGCCGAGGCCT GCATGTCCGGCTGAAGGCTGAGTGTCCGGCTGAGGCCTGAGCGAGTGTCCAGCCAAGGGC TGAGTGTCCAGCACACCTGCCGTCTTCACTTCCCCACAGGCTGGCGCTCGGCTCCACCCC AGGGCCAGCTTTCCTCACCAGGAGCCCGGCTTCCACTCCCCACATAGGAATAGTCCATC CCCAGATTCGCCATTGTTCACCCCTCGCCCTGCCCTCCTTTGCCTTCCACCCCCACCATC CAGGTGGAGACCCTGAGAAGGACCCTGGGAGCTCTGGGAATTTGGAGTGACCAAAGGTGT GCCCTGTACACAGGCGAGGACCCTGCACCTGGATGGGGGTCCCTGTGGGTCAAATTGGGG AAAAAAAAAAAA





MetSerValTyrValValGluLeuLeuArgSerPhePhe TyrValThrGluThrThrPheGlnLysAsnArgLeuPhe PheTyrArgLysSerValTrpSerLysLeuGlnSerIle GlyIleArgGlnHisLeuLysArgValGlnLeuArgGlu LeuSerGluAlaGluValArgGlnHisArgGluAlaArg ProAlaLeuLeuThrSerArgLeuArgPheIleProLys ProAspGlyLeuArgProIleValAsnMetAspTyrVal ValGlyAlaArgThrPheArgArgGluLysArgAlaGlu ArgLeuThrSerArgValLysAlaLeuPheSerValLeu AsnTyrGluArgAlaArgArgProGlyLeuLeuGlyAla SerValLeuGlyLeuAspAspIleHisArgAlaTrpArg GluLeuTyrPheValLysValAspValThrGlyAlaTyr AspThrIleProGlnAspArgLeuThrGluValIleAla SerIleIleLysProGlnAsnThrTyrCysValArgArg TyrAlaValValGlnLysAlaAlaHisGlyHisValArg LysAlaPheLysSerHisValLeuArgProValProGly AspProAlaGlyLeuHisProLeuHisAlaAlaLeuGln ProValLeuArgArgHisGlyGluGlnAlaValCysGly AspSerAlaGlyArgAlaAlaProAlaPheGlyGly

FIG. 19

GCA	GCGC'	rgcg:	rccto	GCTG	CGCA	CGTG	GGAA	GCCC	rggco	CCCG	GCCA	cccc	CGCG	1 met ATG
pro CCG	arg CGC	ala GCT	pro CCC	arg CGC	cys TGC	arg CGA	ala GCC	10 val GTG	arg CGC	ser TCC	leu CTG	leu CTG	arg CGC	ser AGC
											val GTG			
gly GGG	pro CCC	gln CAG	gly GGC	trp TGG	arg CGG	leu CTG	val GTG	40 gln CAG	arg CGC	gly GGG	asp GAC	pro CCG	ala GCG	ala GCT
phe TTC	arg CGC	ala GCG	50 leu CTG	val GTG	ala GCC	gln CAG	cys TGC	leu CTG	val GTG	cys TGC	val GTG	pro CCC	60 trp TGG	asp GAC
											gln CAG			
											leu CTG			
											leu CTG			
											ser AGC			

FIG. 20A





130 tyr leu pro asn thr val thr asp ala leu arg gly ser gly ala TAC CTG CCC AAC ACG GTG ACC GAC GCA CTG CGG GGG AGC GGG GCG 140 trp gly leu leu arg arg val gly asp asp val leu val his TGG GGG CTG CTG CGC CGC GTG GGC GAC GAC GTG CTG GTT CAC 160 leu leu ala arg cys ala leu phe val leu val ala pro ser cys CTG CTG GCA CGC TGC GCG CTC TTT GTG CTG GTG GCT CCC AGC TGC ala tyr gln val cys gly pro pro leu tyr gln leu gly ala ala GCC TAC CAG GTG TGC GGG CCG CCG CTG TAC CAG CTC GGC GCT GCC thr gln ala arg pro pro pro his ala ser gly pro arg arg ACT CAG GCC CGG CCC CCG CCA CAC GCT AGT GGA CCC CGA AGG CGT leu gly cys glu arg ala trp asn his ser val arg glu ala gly CTG GGA TGC GAA CGG GCC TGG AAC CAT AGC GTC AGG GAG GCC GGG val pro leu gly leu pro ala pro gly ala arg arg gly gly GTC CCC CTG GGC CTG CCA GCC CCG GGT GCG AGG AGG CGC GGG GGC 230 240 ser ala ser arg ser leu pro leu pro lys arg pro arg arg gly AGT GCC AGC CGA AGT CTG CCG TTG CCC AAG AGG CCC AGG CGT GGC ala ala pro glu pro glu arg thr pro val gly gln gly ser trp GCT GCC CCT GAG CCG GAG CGG ACG CCC GTT GGG CAG GGG TCC TGG 260 ala his pro gly arg thr arg gly pro ser asp arg gly phe cys GCC CAC CCG GGC AGG ACG CGT GGA CCG AGT GAC CGT GGT TTC TGT 280 val val ser pro ala arg pro ala glu glu ala thr ser leu glu GTG GTG TCA CCT GCC AGA CCC GCC GAA GAA GCC ACC TCT TTG GAG 290 300 gly ala leu ser gly thr arg his ser his pro ser val gly arg GGT GCG CTC TCT GGC ACG CGC CAC TCC CAC CCA TCC GTG GGC CGC gln his his ala gly pro pro ser thr ser arg pro pro arg pro CAG CAC CAC GCG GGC CCC CCA TCC ACA TCG CGG CCA CCA CGT CCC 320 trp asp thr pro cys pro pro val tyr ala glu thr lys his phe TGG GAC ACG CCT TGT CCC CCG GTG TAC GCC GAG ACC AAG CAC TTC

FIG. 20B





340 leu tyr ser ser gly asp lys glu gln leu arg pro ser phe leu CTC TÂC TCC TCA ĞGC GAC AĀG ĞAG CAG CTG CGG CCC TCC TTC CTA 350 leu ser ser leu arg pro ser leu thr gly ala arg arg leu val CTC AGC TCT CTG AGG CCC AGC CTG ACT GGC GCT CGG AGG CTC GTG glu thr ile phe leu gly ser arg pro trp met pro gly thr pro GAG ACC ATC TTT CTG GGT TCC AGG CCC TGG ATG CCA GGG ACT CCC 380 390 arg arg leu pro arg leu pro gln arg tyr trp gln met arg pro CGC AGG TTG CCC CGC CTG CCC CAG CGC TAC TGG CAA ATG CGG CCC leu phe leu glu leu leu gly asn his ala gln cys pro tyr gly CTG TTT CTG GAG CTG CTT GGG AAC CAC GCG CAG TGC CCC TAC GGG 410 val leu leu lys thr his cys pro leu arg ala ala val thr pro GTG CTC CTC AAG ACG CAC TGC CCG CTG CGA GCT GCG GTC ACC CCA 430 ala ala gly val cys ala arg glu lys pro gln gly ser val ala GCA GCC GGT GTC TGT GCC CGG GAG AAG CCC CAG GGC TCT GTG GCG ala pro glu glu glu asp thr asp pro arg arg leu val gln leu GCC CCC GAG GAG GAG GAC ACA GAC CCC CGT CGC CTG GTG CAG CTG leu arg gln his ser ser pro trp gln val tyr gly phe val arg CTC CGC CAG CAC AGC AGC CCC TGG CAG GTG TAC GGC TTC GTG CGG ala cys leu arg arg leu val pro pro gly leu trp gly ser arg GCC TGC CTG CGC CGG CTG GTG CCC CCA GGC CTC TGG GGC TCC AGG 490 his asn glu arg arg phe leu arg asn thr lys lys phe ile ser CAC AAC GAA CGC CGC TTC CTC AGG AAC ACC AAG AAG TTC ATC TCC 500 leu gly lys his ala lys leu ser leu gln glu leu thr trp lys CTG GGG AAG CAT GCC AAG CTC TCG CTG CAG GAG CTG ACG TGG AAG 520 met ser val arg asp cys ala trp leu arg arg ser pro gly val ATG AGC GTG CGG GAC TGC GCT TGG CTG CGC AGG AGC CCA GGG GTT gly cys val pro ala ala glu his arg leu arg glu glu ile leu GGC TGT GTT CCG GCC GCA GAG CAC CGT CTG CGT GAG GAG ATC CTG

FIG. 20C





550 ala lys phe leu his trp leu met ser val tyr val val glu leu GCC AAG TTC CTG CAC TGG CTG ATG AGT GTG TAC GTC GAG CTG 560 leu arg ser phe phe tyr val thr glu thr thr phe gln lys asn CTC AGG TCT TTC TTT TAT GTC ACG GAG ACC ACG TTT CAA AAG AAC 580 arg leu phe phe tyr arg lys ser val trp ser lys leu gln ser AGG CTC TTT TTC TAC CGG AAG AGT GTC TGG AGC AAG TTG CAA AGC ile gly ile arg gln his leu lys arg val gln leu arg glu leu ATT GGA ATC AGA CAG CAC TTG AAG AGG GTG CAG CTG CGG GAG CTG ser glu ala glu val arg gln his arg glu ala arg pro ala leu TCG GAA GCA GAG GTC AGG CAG CAT CGG GAA GCC AGG CCC GCC CTG 620 leu thr ser arg leu arg phe ile pro lys pro asp gly leu arg CTG ACG TCC AGA CTC CGC TTC ATC CCC AAG CCT GAC GGG CTG CGG 640 pro ile val asn met asp tyr val val gly ala arg thr phe arg CCG ATT GTG AAC ATG GAC TAC GTC GTG GGA GCC AGA ACG TTC CGC arg glu lys arg ala glu arg leu thr ser arg val lys ala leu AGA GAA AAG AGG GCC GAG CGT CTC ACC TCG AGG GTG AAG GCA CTG 670 phe ser val leu asn tyr glu arg ala arg arg pro gly leu leu TTC AGC GTG CTC AAC TAC GAG CGG GCG CGC CCC GGC CTC CTG gly ala ser val leu gly leu asp asp ile his arg ala trp arg GGC GCC TCT GTG CTG GGC CTG GAC GAT ATC CAC AGG GCC TGG CGC 700 thr phe val leu arg val arg ala gln asp pro pro pro glu leu ACC TTC GTG CTG CGT GTG CGG GCC CAG GAC CCG CCT GAG CTG 710 tyr phe val lys val asp val thr gly ala tyr asp thr ile pro TAC TTT GTC AAG GTG GAT GTG ACG GGC GCG TAC GAC ACC ATC CCC 730 gln asp arg leu thr glu val ile ala ser ile ile lys pro gln CAG GAC AGG CTC ACG GAG GTC ATC GCC AGC ATC ATC AAA CCC CAG asn thr tyr cys val arg arg tyr ala val val gln lys ala ala AAC ACG TAC TGC GTG CGT CGG TAT GCC GTG GTC CAG AAG GCC GCC

FIG. 20D



760



his gly his val arg lys ala phe lys ser his val leu arg pro CAT GGG CAC GTC CGC AAG GCC TTC AAG AGC CAC GTC CTA CGT CCA 780 val pro gly asp pro ala gly leu his pro leu his ala ala leu GTG CCA GGG GAT CCC GCA GGG CTC CAT CCT CTC CAC GCT GCT CTG 790 gln pro val leu arg arg his gly glu gln ala val cys gly asp CAG CCT GTG CTA CGG CGA CAT GGA GAA CAA GCT GTT TGC GGG GAT ser ala gly arg ala ala pro ala phe gly gly OP TCG GCG GGA CGG GCT GCT CCT GCG TTT GGT GGA TGA TTTCTTGTTGGT GACACCTCACCTCACCCACGCGAAAACCTTCCTCAGGACCCTGGTCCGAGGTGTCCCTGA GTATGGCTGCTGTGAACTTGCGGAAGACAGTGGTGAACTTCCCTGTAGAAGACGAGGC CCTGGGTGGCACGGCTTTTGTTCAGATGCCGGCCCACGGCCTATTCCCCTGGTGCGGCCT GCTGCTGGATACCCGGACCCTGGAGGTGCAGAGCGACTACTCCAGCTATGCCCGGACCTC CATCAGAGCCAGTCTCACCTTCAACCGCGGCTTCAAGGCTGGGAGGAACATGCGTCGCAA ACTCTTTGGGGTCTTGCGGCTGAAGTGTCACAGCCTGTTTCTGGATTTGCAGGTGAACAG CCTCCAGACGGTGTGCACCAACATCTACAAGATCCTCCTGCTGCAGGCGTACAGGTTTCA CGCATGTGTGCTGCAGCTCCCATTTCATCAGCAAGTTTGGAAGAACCCCACATTTTTCCT GCGCGTCATCTCTGACACGGCCTCCCTCTGCTACTCCATCCTGAAAGCCAAGAACGCAGG GATGTCGCTGGGGGCCAAGGGCGCCGCCGCCTCTGCCCTCCGAGGCCGTGCAGTGGCT GTGCCACCAAGCATTCCTGCTCAAGCTGACTCGACACCGTGTCACCTACGTGCCACTCCT GGGGTCACTCAGGACAGCCCAGACGCAGCTGAGTCGGAAGCTCCCGGGGACGACGCTGAC TGCCCTGGAGGCCGCGCCAACCCGGCACTGCCCTCAGACTTCAAGACCATCCTGGACTG CTACGTCCCAGGGGGGGGGGGCCCACACCCAGGCCCGCACCGCTGGGAGTCTGAGG CCTGAGTGAGTGTTTGGCCGAGGCCTGCATGTCCGGCTGAAGGCTGAGTGTCCGGCTGAG GCCTGAGCGAGTGTCCAGCCAAGGGCTGAGTGTCCAGCACACCTGCCGTCTTCACTTCCC CACAGGCTGGCGCTCCACCCCAGGGCCAGCTTTTCCTCACCAGGAGCCCGGCTTC CACTCCCCACATAGGAATAGTCCATCCCCAGATTCGCCATTGTTCACCCCTCGCCCTGCC CTCCTTTGCCTTCCACCCCCACCATCCAGGTGGAGACCCTGAGAAGGACCCTGGGAGCTC TGGGAATTTGGAGTGACCAAAGGTGTGCCCTGTACACAGGCGAGGACCCTGCACCTGGAT GGGGGTCCCTGTGGGTCAAATTGGGGGGAGGTGCTGTGGGAGTAAAATACTGAATATATG

FIG. 20E



1 CCATGGGACCCACTGCAGGGCAGCTGGGAGGCTGCAGGCTTCAGGTCCCAGTGGGGTTG GGTACCCTGGGTGACGTCCCCGTCGACCCTCCGACGTCCGAAGTCCAGGGTCACCCCAAC 61 CCATCTGCCAGTAGAAACCTGATGTAGAATCAGGGCGCGAGTGTGGACACTGTCCTGAAT GGTAGACGGTCATCTTTGGACTACATCTTAGTCCCGCGCTCACACCTGTGACAGGACTTA 121 CTCAATGTCTCAGTGTGTGCTGAAACATGTAGAAATTAAAGTCCATCCCTCCTACTCTAC GAGTTACAGAGTCACACGACTTTGTACATCTTTAATTTCAGGTAGGGAGGATGAGATG 181 TGGGATTGAGCCCCTTCCCTATCCCCCCCAGGGGCAGAGGAGTTCCTCTCACTCCTGTG ACCCTAACTCGGGGAAGGGATAGGGGGGGGTCCCCGTCTCCTCAAGGAGAGTGAGGACAC 241 GAGGAAGGAATGATACTTTGTTATTTTTCACTGCTGGTACTGAATCCACTGTTTCATTTG CTCCTTCCTTACTATGAAACAATAAAAGTGACGACCATGACTTAGGTGACAAAGTAAAC ********** ****************** 361 TGCAATGGCGCGATCTTGGCTTACTGCAGCCTCTGCCTCCCAGGTTCAAGTGATTCTCCT ACGTTACCGCGCTAGAACCGAATGACGTCGGAGACGGAGGGTCCAAGTTCACTAAGAGGA alu ***************** 421 GCTTCCGCCTCCCATTTGGCTGGGATTACAGGCACCCGCCACCATGCCCAGCTAATTTTT CGAAGGCGGAGGTAAACCGACCCTAATGTCCGTGGGCGGTGGTACGGGTCGATTAAAAA ******************* 481 TGTATTTTAGTAGAGACGGGGGTGGGGGTTCACCATGTTGGCCAGGCTGGTCTC ACATAAAAATCATCTCTGCCCCCACCCCCACCCCAAGTGGTACAACCGGTCCGACCAGAG CAP =======> ********************* 541 GAACTTCTGACCTCAGATGATCCACCTGCCTCTGCCTCCTAAAGTGCTGGGATTACAGGT CTTGAAGACTGGAGTCTACTAGGTGGACGGAGACGGAGGATTTCACGACCCTAATGTCCA ******* 601 GTGAGCCACCATGCCCAGCTCAGAATTTACTCTGTTTAGAAACATCTGGGTCTGAGGTAG CACTCGGTGGTACGGGTCGAGTCTTAAATGAGACAAATCTTTGTAGACCCAGACTCCATC CCAAT ******* 661 GAAGCTCACCCCACTCAAGTGTTGTGGTGTTTTAAGCCAATGATAGAATTTTTTTATTGT 721 TGTTAGAACACTCTTGATGTTTTACACTGTGATGACTAAGACATCATCAGCTTTTCAAAG ACAATCTTGTGAGAACTACAAAATGTGACACTACTGATTCTGTAGTAGTCGAAAAGTTTC

FIG. 21A



	CAP
781	************************** ACACACTAACTGCACCCATAATACTGGGGTGTCTTCTTGGGTATCAGCGATCTTCATTGAA TGTGTGATTGACGTGGGTATTATGACCCCACAGAAGACCCATAGTCGCTAGAAGTAACTT
	CAP

841	TGCCGGGAGGCGTTTCCTCGCCATGCACATGGTGTTAATTACTCCAGCATAATCTTCTGC ACGGCCCTCCGCAAAGGAGCGGTACGTGTACCACAATTAATGAGGTCGTATTAGAAGACG
	***>
901	TTCCATTTCTTCTCTCTCTCTTTTAAAATTGTGTTTTCTATGTTGGCTTCTCTGCAGAG AAGGTAAAGAAGAGAAG
	CAP

961	AACCAGTGTAAGCTACAACTTAACTTTTGTTGGAACAAATTTTCCAAACCGCCCCTTTGC TTGGTCACATTCGATGTTGAATTGAA
1021	${\tt CCTAGTGGCAGAGACAATTCACAAACACAGCCCTTTAAAAAGGCTTAGGGATCACTAAGGGGATCACCGAATCCCTAGTGATTCCGGGGAAATTTTTCCGAATCCCTAGTGATTCCGGGGAAATTTTTCCGAATCCCTAGTGATTCCGGGGAAATTTTTCCGGAATCCCTAGTGATTCCGGGGAAATTTTTCCGGAATCCCTAGTGATTCCGGGGAAATTTTTCCGGAATCCCTAGTGATTCCGGGGAAATTTTTCCGGAATCCCTAGTGATTCCGGGGAAATTTTTCCGGAATCCCTAGTGATTCCGGGGAAATTTTTCCGGAATCCCTAGTGATTCCGGGGAAATTTTTCCGGAATCCCTAGTGATTCCCGAATCCCTAGTGATTCCGGGGAAATTTTTCCGGAATCCCTAGTGATTCCCGAATCCCTAGTGATTCCCTAGTGATTCCCGAATCCCTAGTGATTCCCGAATCCCTAGTGATTCCCGAATCCCTAGTGATTCCCTAGTGATTCCCGAATCCCTAGTGATTCCCGAATCCCTAGTGATTCCCGAATCCCTAGTGATTCCCTAGTGATTCCCGAATCCCTAGTGATTCCTAGTGATTCCTAGTGATTCCTAGTGATTCCTAGTGATTCCTAGTGATTCCTAGTGATTCCTAGTGATTCCTAGTGATTCTAGTGATTCTAGTGATTCTAGTGATTCTAGTGATTCTAGTGATTCTAGTGATTCTAGTGATTCTAGTGATTCTAGTGATTTTTTTT$
1081	GGATTTCTAGAAGAGCGACCCGTAATCCTTAAGTATTTACAAGACGAGGCTAACCTCCAG
	$\tt CCTAAAGATCTTCTCGCTGGGCATTAGGAATTCATAAATGTTCTGCTCCGATTGGAGGTCCGGATTGGAGGTCCGGATTGGAGGTCGGAGGAGGTCGGAGGAGGTCGGAGGAGGAGGAGGAGGAGGAGGAGGAGGAGGAGGAGGA$
1141	CGAGCGTGACAGCCCAGGGAGGGTGCGAGGCCTGTTCAAATGCTAAGCTTCCATAAATAA
	GCTCGCACTGTCGGGTCCCTCCCACGCTCCGGACAAGTTTACGATTCGAAGGTATTTATT
1201	AGCAAATTTCCTCCGGCAGTTTCTGGAAAGTAGGAAAGGTTAACATTTAAGGTTGCGTTT
	TCGTTTAAAGGAGGCCGTCAAAGACCTTTCATCCTTTCCAATTGTAAATTCCAACGCAAA
1261	GTTAGCATTTCAGTGTTTGCCGACCTCAGCTAACAGCATCCCTGCAAGGCCTCGGGAGACCAATCGTAAAGTCACAAACGGCTGGAGTCGATTGTCGTAGGGACGTTCCGGAGCCCTCTG
1321	CCAGAAGTTTCTCGCCCCTTAGATCCAAACTTGAGCAACCCGGAGTCTGGATTCCTGGGA
	GGTCTTCAAAGAGCGGGAATCTAGGTTTGAACTCGTTGGGCCTCAGACCTAAGGACCCT
	TopoII

1381	AGTCCTCAGCTGTCCTGCGGTTGTGCCGGGGCCCCAGGTCTGGAGGGGGACCAGTGGCCGT TCAGGAGTCGACAGGACGCCAACACGGCCCCGGGGTCCAGACCTCCCCTGGTCACCGGCA
1441	
	CACCGAAGATGACGACCCGACCTTCAGCCCGGAGGATCGAGACGTCAGGCTCCGAACCTC
1501	CCAGGTGCCTGGACCCCGAGGCTGCCCTCCACCCTGTGCGGGCGG
	GGTCCACGGACCTGGGGCTCCGACGGGGGGGGGGGCCCCCCCC
1561	TGGCCTCATCTGCCAGACAGAGTGCCGGGGCCCAGGGTCAAGGCCGTTGTGGCTGGTGTG
	ACCGGAGTAGACGGTCTCTCACGGCCCCGGGTCCCAGTTCCGGCAACACCGACCAC
1621	${\tt AGGCGCCCGGTGCGCCAGCAGCAGCAGCAGCGCCTTGCCCACCCTTTCTCGACGG}$
	TCCGCGGGCCACGCCGGTCGTCCTCGCGGACCGAGGTAAAGGGTGGGAAAGAGCTGCC

FIG. 21B



- 1681 GACCGCCCGGTGGTGATTAACAGATATTGGGGTGGTTTGCTCATGGTGGGGACCCCTT CTGGCGGGGCCACCCACTAATTGTCTATAACCCCACCAAACGAGTACCACCCCTGGGGAA
- 1801 GGAAGTGTTGCAGGGAGGCACTCCGGGAGGTCCCGCGTGCCGTCCAGGGAGCAATGCGT CCTTCACAACGTCCCTCCGTGAGGCCCTCCAGGGCGCACGGCAGGTCCCTCGTTACGCA
- 1861 CCTCGGGTTCGTCCCCAGCCGCGTCTACGCGCCTCCGTCCTCCCCTTCACGTCCGGCATT
 GGAGCCCAAGCAGGGGTCGGCGCAGATGCGCGGAGGCAGGAGGGGAAGTGCAGGCCGTAA
- 1921 CGTGGTGCCCGGAGCCCGACGCCCGCGTCCGGACCTGGAGCCCTGGGTCTCCGGA GCACCACGGGCCTCGGGCTGCGGGCCCTGGACCTCCGTCGGGACCCAGAGGCCT
- 1981 TCAGGCCAGCGCCAAAGGGTCGCCGCACCTGTTCCCAGGGCCTCCACATCATGGC AGTCCGGTCGCCGGTTTCCCAGCGGCGTGCGTGGACAAGGGTCCCGGAGGTGTAGTACCG

Sp1

- 2161 CCCCGGGTCCGCCCGGAGCAGCTGCGCTGTCGGGGCCAGGCCGGGCTCCCAGTGGATTCG
 GGGGCCCAGGCGGGCCTCGTCGACGCGACAGCCCCGGTCCGGCCCGAGGGTCACCTAAGC
- 2221 CGGGCAACAGACGCCCAGGACCGCGTTCCCACGTGGCGAGGGACTGGGGACCCGGGCA GCCCGTTGTCTGCGGGTCCTGGCGCGAAGGGTGCACCGCCTCCCTGACCCCTGGGCCCGT

Sp1

E2F

- 2341 GAACCCTTCCCGGGTCCCGGCCCAGCCCTTCCGGGCCATCCCAGCCCGTCCCGTTCCT CTTGGGAAGGCCCAGGGCCGGGTCGGGAAGGCCCGGTAGGGTCGGGCAAGGA

Sp1

E2F NFkB

2401 TTTCCGCGGCCCCCCCTCTCCTCGCGGCGCGAGTTTCAGGCAGCGCTGCGTCCTGCTGC
AAAGGCGCCGGGGGGGAGAGGAGCGCCGCGCTCAAAGTCCGTCGCGACGCAGGACGACG

hTRT5' *********************

- 2521 CCGTGCGCTGCTGCGCAGCCACTACCGCGAGGTGCTGCCGCTGGCCACGTTCGTGC
 GGCACGCGAGGGACGCGTCGGTGATGGCGCTCCACGACGCGACCGCTGCAAGCACG

FIG. 21C

E2F



2581 GGCGCCTGGGGCCCCAGGGCTGGCGGCTGCTGCAGCGCGGGGACCCGGCGGCTTTCCGCG CCGCGGACCCCGGGGTCCCGACCGCCGACCGCGCCCCTGGGCCGCCGAAAGGCGC NFkB ======== ************* 2701 CCTCCTTCCGCCAGGTGGGCCTCCCCGGGGTCGGCGTCCGGCTGGGGTTGAGGGCGGCCG GGAGGAAGGCGGTCCACCCGGAGGGGCCCCAGCCGCAGCCGACCCCAACTCCCGCCGGC Topo II cleavag NFkB ++++++++ NFkB _____ Intron1 ********************* 2761 GGGGGAACCAGCGACATGCGGAGAGCAGCGCAGGCGACTCAGGGCGCTTCCCCCGCAGGT CCCCTTGGTCGCTGTACGCCTCTCGTCGCGTCGCTGAGTCCCGCGAAGGGGGGCGTCCA e site :::: CAGGACGGACTTCCTCGACCACCGGGCTCACGACGTCTCCGACACGCTCGCGCCGCTT 2881 GAACGTGCTGGCCTTCGGCTTCGCGCTGGACGGGGCCCGCGGGGGCCCCCCGAGGC 2941 CTTCACCACCAGCGTGCGCAGCTACCTGCCCAACACGGTGACCGACGCACTGCGGGGGAG GAAGTGGTGGTCGCACGCGTCGATGGACGGGTTGTGCCACTGGCTGCGTGACGCCCCCTC 3001 CGGGGCGTGGGGCTGCTGCTGCGCCGCGTGGGCGACGACGTGCTGGTTCACCTGCTGGC GCCCGCACCCCGACGACGACGCGCGCGCACCCGCTGCTGCACGACCAAGTGGACGACCA 3061 ACGCTGCGCGCTCTTTGTGCTGGTGGCTCCCAGCTGCGCCTACCAGGTGTGCGGGCCGCC TGCGACGCGCGAGAAACACGACCACCGAGGGTCGACGCGGGATGGTCCACACGCCCGGCGG

3181 AAGGCGTCTGGGATGCGAACGGGCCTGGAACCATAGCGTCAGGGAGGCCGGGGTCCCCCT TTCCGCAGACCCTACGCTTGCCCGGACCTTGGTATCGCAGTCCCTCCGGCCCCAGGGGGA

3301 GCCCAAGAGGCCCAGGCGTGGCGCTGCCCCTGAGCCGGAGCGGACGCCCGTTGGGCAGGG CGGGTTCTCCGGGTCCGCACCGCGACGGGGACTCGGCCTCGCCTGCGGGCAACCCGTCCC

FIG. 21D



3361	GTCCTGGGCCCACCCGGGCAGGACGCGTGGACCGAGTGACCGTGGTTTCTGTGTGTG
3421	ACCTGCCAGACCCGCCGAAGAAGCCACCTCTTTGGAGGGTGCGCTCTCTGGCACGCGCCATGGACGGTCTGGGCGGCTTCTTCGGTGGAGAAACCTCCCACGCGAGAGACCGTGCGCGGT
3481	CTCCCACCCATCCGTGGGCCGCCAGCACCACGCGGGCCCCCATCCACATCGCGGCCACCGAGGGTGGGT
3541	ACGTCCCTGGGACACGCCTTGTCCCCCGGTGTACGCCGAGACCAAGCACTTCCTCTACTCTGCAGGGACCCTGTGCGGAACAGGGGGCCACATGCGGCTCTGGTTCGTGAAGGAGATGAG
3601	CTCAGGCGACAAGGAGCAGCTGCGGCCCTCCTTCCTACTCAGCTCTCTGAGGCCCAGCCTGAGTCCGCTGTTCCTCGTCGACGCCGGGAGGAAGGA
3661	GACTGGCGCTCGGAGGCTCGTGGAGACCATCTTTCTGGGTTCCAGGCCCTGGATGCCAGGCTGACCGCGAGCCTCCGAGCACCTCTGGTAGAAAGACCCAAGGTCCGGGACCTACGGTCC
3721	GACTCCCGCAGGTTGCCCCGCCTGCCCCAGCGCTACTGGCAAATGCGGCCCCTGTTTCTCTGAGGGGCGTCCAACGGGGCGGACGGGGTCGCGATGACCGTTTACGCCGGGGACAAAGA
3781	GGAGCTGCTTGGGAACCACGCGCAGTGCCCCTACGGGGTGCTCCTCAAGACGCACTGCCCCCTCGACGAACCCTTGGTGCGCGTCACGGGGATGCCCCACGAGGAGTTCTGCGTGACGGG
3841	GCTGCGAGCTGCGGTCACCCCAGCAGCCGGTGTCTGTGCCCGGGAGAAGCCCCAGGGCTCCCGACGCTCGACGCCAGTGGGGTCGTCGGCCACAGACACGGGCCCTCTTCGGGGTCCCGAG
3901	TGTGGCGGCCCCGAGGAGGAGGACACAGACCCCCGTCGCCTGGTGCAGCTGCTCCGCCAACACGCCGGGGGCTCCTCCTCCTGTGTCTGGGGGCAGCGGACCACGTCGACGAGGGGGT
3961	GCACAGCAGCCCTGGCAGGTGTACGGCTTCGTGCGGGGCCTGCCT
4021	CCCAGGCCTCTGGGGCTCCAGGCACAACGAACGCCGCTTCCTCAGGAACACCAAGAAGTT GGGTCCGGAGACCCCGAGGTCCGTGTTGCTTGCGGCGAAGGAGTCCTTGTGGTTCTTCAA
4081	CATCTCCCTGGGGAAGCATGCCAAGCTCTCGCTGCAGGAGCTGACGTGGAAGATGAGCGT GTAGAGGGACCCCTTCGTACGGTTCGAGAGCGACGTCCTCGACTGCACCTTCTACTCGCA
4141	**************************************
4201	Intron2 ***********************************
4261	**************************************
4321	****> GTGATCGAGGTCGAC CACTAGGTGGAGGTCGAC

FIG. 21E



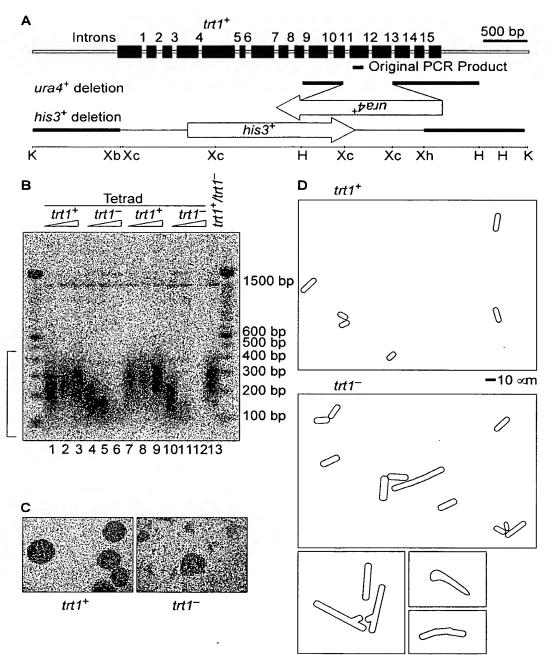


FIG. 22



FIG. 23

TCTACCTTGACAGACCTCCAGCCGTACATGCGACAGTTCGTGGCTCACCTGCAGGAG ACCAGCCGCTGAGGGATGCCGTCGTCATCGAGCAGAGCTCCTCCCTGAATGAGGCC AGCAGTGGCCTCTTCGACGTCTTCCTACGCTTCATGTGCCACCACGCCGTGCGCATC AGGGGCAAGTC

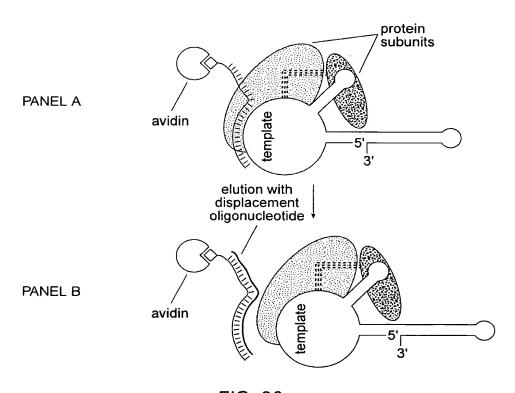
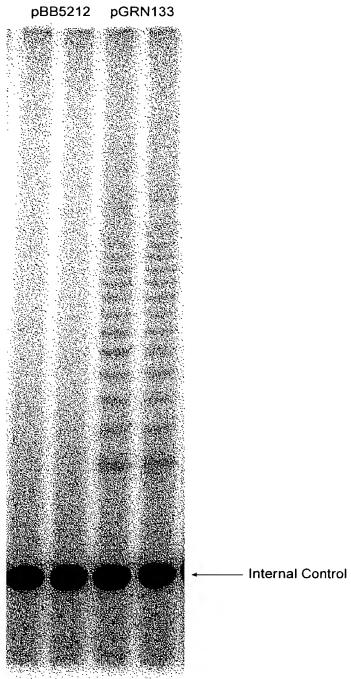


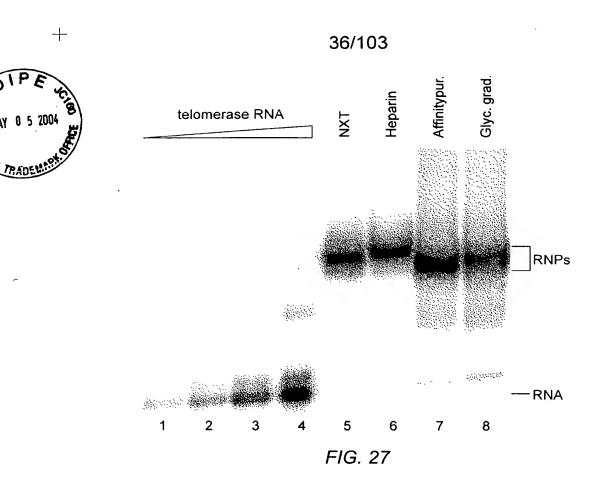
FIG. 26





Approximate Cell No. 5,000 5,000 5,000 5,000

FIG. 25



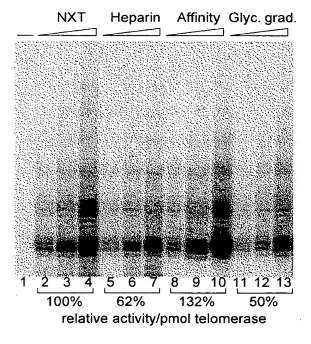
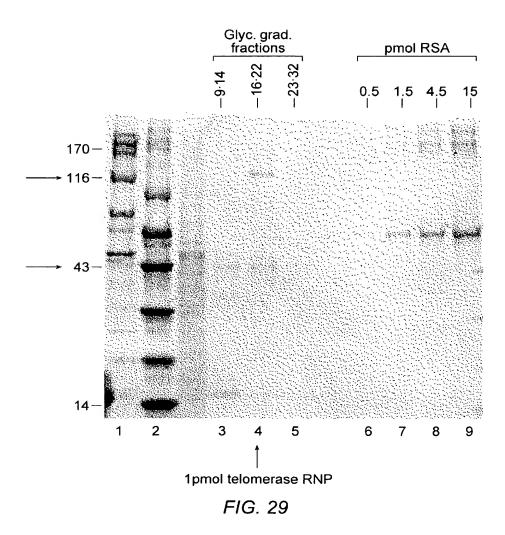


FIG. 28







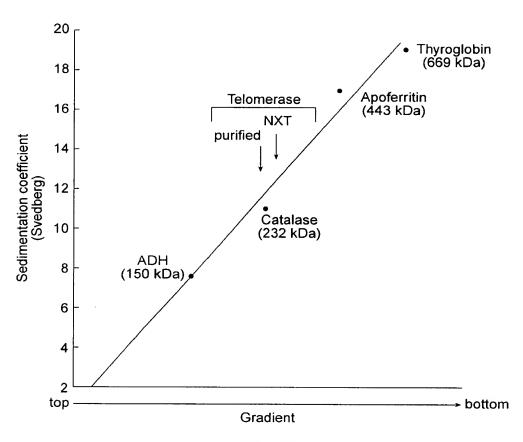


FIG. 30

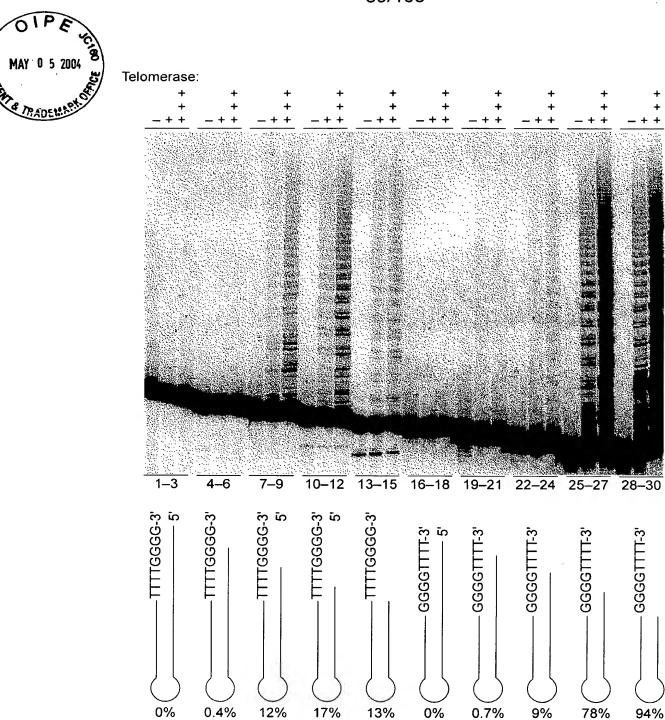


FIG. 31

% primer extended

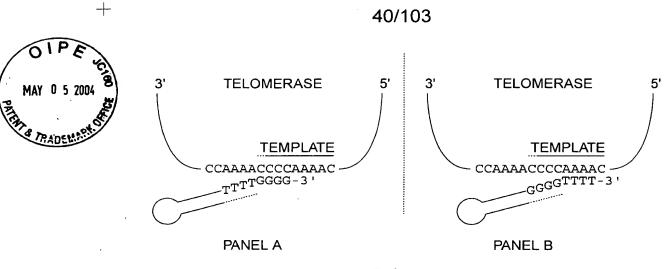


FIG. 32

1	CCCCAAAACC	CCAAAACCCC	AAAACCCCTA	TAAAAAAAGA	AAAAATTGAG
51	GTAGTTTAGA	AATAAAATAT	TATTCCCGCA	CAAATGGAGA	TGGATATTGA
101	TTTGGATGAT	ATAGAAAATT	TACTTCCTAA	TACATTCAAC	AAGTATAGCA
151	GCTCTTGTAG	TGACAAGAAA	GGATGCAAAA	CATTGAAATC	TGGCTCGAAA
201	TCGCCTTCAT	TGACTATTCC	AAAGTTGCAA	AAACAATTAG	AGTTCTACTT
251	CTCGGATGCA	AATCTTTATA	ACGATTCTTT	CTTGAGAAAA	TTAGTTTTAA
301	AAAGCGGAGA	GCAAAGAGTA	GAAATTGAAA	CATTACTAAT	GTTTAAATAA
351	AATCAGGTAA	TGAGGATTAT	TCTATTTTTT	AGATCACTTC	TTAAGGAGCA
401	TTATGGAGAA	AATTACTTAA	TACTAAAAGG	TAAACAGTTT	GGATTATTTC
451	CCTAGCCAAC	AATGATGAGT	ATATTAAATT	CATATGAGAA	TGAGTCAAAG
501	GATCTCGATA	CATCAGACTT	ACCAAAGACA	AACTCGCTAT	AAAACGCAAG
551	AAAAAGTTTG	ATAATCGAAC	AGCAGAAGAA	CTTATTGCAT	TTACTATTCG
601	TATGGGTTTT	ATTACAATTG	TTTTAGGTAT	CGACGGTGAA	CTCCCGAGTC
651	TTGAGACAAT	TGAAAAAGCT	GTTTACAACT	GAAGGAATCG	CAGTTCTGAA
701	AGTTCTGATG	TGTATGCCAT	TATTTTGTGA	ATTAATCTCA	AATATCTTAT
751	CTCAATTTAA	TGGATAGCTA	TAGAAACAAA	CCAAATAAAC	CATGCAAGTT
801	TAATGGAATA	TACGTTAAAT	CCTTTGGGAC	AAATGCACAC	TGAATTTATA
851	TTGGATTCTT	AAAGCATAGA	TACACAGAAT	GCTTTAGAGA	CTGATTTAGC
901	TTACAACAGA	TTACCTGTTT	TGATTACTCT	TGCTCATCTC	TTATATCTTT
951	AAAAGAAGCA	GGCGAAATGA	AAAGAAGACT	AAAGAAAGAG	ATTTCAAAAT
1001	TTGTTGATTC	TTCTGTAACC	GGAATTAACA	ACAAGAATAT	TAGCAACGAA
1051	AAAGAAGAAG	AGCTATCACA	ATCCTGATTC	TTAAAGATTT	CAAAAATTCC
1101	AGGTAAGAGA	GATACATTCA	TTAAAATTCA	TATATTATAG	TTTTTCATTT
1151	CACAGCTGTT	ATTTTCTTTT	ATCTTAACAA	TATTTTTTGA	TTAGCTGGAA
1201	GTAAAAAGTA	TCAAATAAGA	GAAGCGCTAG	ACTGAGGTAA	CTTAGCTTAT
1251	TCACATTCAT	AGATCGACCT	TCATATATCC	AATACGATGA	TAAGGAAACA
1301	GCAGTCATCC	GTTTTAAAAA	TAGTGCTATG	AGGACTAAAT	TTTTAGAGTC
1351	AAGAAATGGA	GCCGAAATCT	TAATCAAAAA	GAATTGCGTC	GATATTGCAA
1401	AAGAATCGAA	CTCTAAATCT	TTCGTTAATA	AGTATTACCA	ATCTTGATTG
1451	ATTGAAGAGA	TTGACGAGGC	AACTGCACAG	AAGATCATTA	AAGAAATAAA
1501	GTAACTTTTA	TTAATTAGAG	AATAAACTAA	ATTACTAATA	TAGAGATCAG
1551	CGATCTTCAA	TTGACGAAAT	AAAAGCTGAA	CTAAAGTTAG	ACAATAAAAA
1601	ATACAAACCT	TGGTCAAAAT	ATTGAGGAAG	GAAAAGAAGA	CCAGTTAGCA
1651	AAAGAAAAA	TAAGGCAATA	AATAAAATGA	GTACAGAAGT	GAAGAAATAA
1701	AAGATTTATT		ATTTATTGAA	AAGAGGGGTT	TTGGGGTTTT
1751	GGGGTTTTGG	GG			

FIG. 34



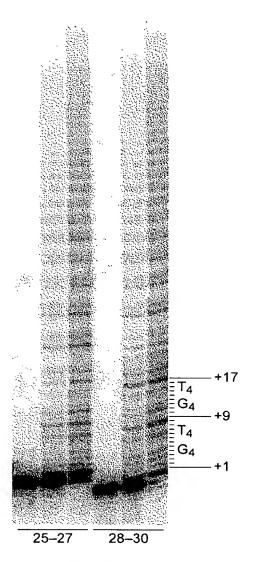


FIG. 33



CCCCAAAACCCCAAAACCCCAAAACCCCTATAAAAAAAGAAAAATTGAGGTAGTTTAGA 1+++ 60 GGGGTTTTGGGGTTTTGGGGATATTTTTTTTTTTTTTT	
a PQNPKTPKPL * KKKKLR * FR - b PKTPKPQNPYKKRKNCGSLE - c PKPQNPKTPIKKEKIEVV * K-	
AATAAAATATTATTCCCGCACAAATGGAGATGGATATTGATTTGGATGATATAGAAAATT 61++ 12 TTATTTTATAATAAGGGCGTGTTTACCTCTACCTATAACTAAACCTACTATATCTTTTAA	0
a N K I L F P H K W R W I L I W M I * K I - b I K Y Y S R T N G D G Y C F G C Y R K F - c * N I I P A Q M E M D I D L D D I E N L -	
TACTTCCTAATACATTCAACAAGTATAGCAGCTCTTGTAGTGACAAGAAAGGATGCAAAA 121+ 18 ATGAAGGATTATGTAAGTTGTTCATATCGTCGAGAACATCACTGTTCTTTCCTACGTTTT	0
a Y F L I H S T S I A A L V V T R K D A K - b T S * Y I Q Q V * Q L L * C Q E R M Q N - c L P N T F N K Y S S S C S D K K G C K T -	
CATTGAAATCTGGCTCGAAATCGCCTTCATTGACTATTCCAAAGTTGCAAAAACAATTAG 181++ 24 GTAACTTTAGACCGAGCTTTAGCGGAAGTAACTGATAAGGTTTCAACGTTTTTGTTAATC	0
a H C N L A R N R L H C L F Q S C K N N * - b I E I W L E I A F I D Y S K V A K T I R - c L K S G S K S P S L T I P K L Q K Q L E -	
AGTTCTACTTCTCGGATGCAAATCTTTATAACGATTCTTTCT	0
a S S T S R M Q I F I T I L S C E N * F * - b V L L L G C K S L * R F F L E K I S F K - c F Y F S D A N L Y N D S F L R K L V L K -	
AAAGCGGAGAGCAAAGAGTAGAAATTGAAACATTACTAATGTTTAAATAAA	0
a	
TGAGGATTATTCTATTTTTTAGATCACTTCTTAAGGAGCATTATGGAGAAAATTACTTAA 361+ 42 ACTCCTAATAAGATAAAAAATCTAGTGAAGAATTCCTCGTAATACCTCTTTTAATGAATT	0
a C G L F Y F L D H F L R S I M E K I T * - b E D Y S I F * I T S * G A L W R K L L N - c R I I L F F R S L L K E H Y G E N Y L I -	

FIG. 35A



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TACTAAAAGGTAAACAGTTTGGATTATTTCCCTAGCCAACAATGATGAGTATATTAAATT 421 ---------- 480 ATGATTTTCCATTTGTCAAACCTAATAAAGGGATCGGTTGTTACTACTCATATAATTTAA Y * K V N S L D Y F P S Q Y * K V N S L D Y F P S Q Q C C V Y * I -T K R * T V W I I S L A N N D E Y I K F b LKGKOFGLFP* PTMMSILNS-CATATGAGAATGAGTCAAAGGATCTCGATACATCAGACTTACCAAAGACAAACTCGCTAT 481 -----+ 540 GTATACTCTTACTCAGTTTCCTAGAGCTATGTAGTCTGAATGGTTTCTGTTTGAGCGATA ${\tt AAAACGCAAGAAAAAGTTTGATAATCGAACAGCAGAAGAACTTATTGCATTTACTATTCG}$ 541 -----+ 600 TTTTGCGTTCTTTTCAAACTATTAGCTTGTCGTCTTCTTGAATAACGTAAATGATAAGC CTQEKVC*SNSRRTYCIYYS KRKKFDNRTAEELIAFTIR b N A R K S L I I E Q Q K N L L H L L F V - ${\tt TATGGGTTTTATTACAATTGTTTTAGGTATCGACGGTGAACTCCCGAGTCTTGAGACAAT}$ ATACCCAAAATAATGTTAACAAAATCCATAGCTGCCACTTGAGGGCTCAGAACTCTGTTA YGFYYNCFRYRRCTPESCDN M G F I T I V L G I D G E L P S L E T I - W V L L Q L F * V S T V N S R V L R Q L b TGAAAAAGCTGTTTACAACTGAAGGAATCGCAGTTCTGAAAGTTCTGATGTGTATGCCAT ACTTTTTCGACAAATGTTGACTTCCTTAGCGTCAAGACTTTCAAGACTACACATACGGTA CKSCLQLKESQFCKFCCVCH -EKAVYNCRNRSSESSDVYAI -KKLFTTEGIAVLKVLMCMPL-TATTTTGTGAATTAATCTCAAATATCTTATCTCAATTTAATGGATAGCTATAGAAACAAA 721 -----+ 780 ATAAAACACTTAATTAGAGTTTATAGAATAGAGTTAAATTACCTATCGATATCTTTGTTT Y F V N * S Q I S Y L N L M D S Y R N K - I L C I N L K Y L I S I * W I A I E T N - F C E L I S N I L S Q F N G * L * K Q T -CCAAATAAACCATGCAAGTTTAATGGAATATACGTTAAATCCTTTGGGACAAATGCACAC GGTTTATTTGGTACGTTCAAATTACCTTATATGCAATTTAGGAAACCCTGTTTACGTGTG P N K P C K F N G I Y V K S F G T N A H - Q I N H A S L M E Y T L N P L G Q M H T - K * T M Q V * W N I R * I L W D K C T L -TGAATTTATATTGGATTCTTAAAGCATAGATACACAGAATGCTTTAGAGACTGATTTAGC ACTTAAATATAACCTAAGAATTTCGTATCTATGTGTCTTACGAAATCTCTGACTAAATCG GFLKHRYTECFRDCF E F I L D S * S I D T Q N A L E T D L A - N L Y W I L K A * I H R M L * R L I * L -

FIG. 35B



TTACAACAGATTACCTGTTTTGATTACTCTTGCTCATCTCTTATATCTTTAAAAGAAGCA
AATGTTGTCTAATGGACAAAACTAATGAGAACGAGTAGAGAATATAGAAATTTTCTTCGT
a L Q Q I T C F D Y S C S S L I S L K E A - b Y N R L P V L I T L A H L L Y L * K K Q - c T T D Y L F C L L L I S Y I F K R S R -
GGCGAAATGAAAAGAAGACTAAAGAAAGAGATTTCAAAATTTGTTGATTCTTCTGTAACC 961+ 1020 CCGCTTTACTTTCTTCTGATTTCTTCTCTAAAGTTTTAAACAACTAAGAAGACATTGG
a G E M K R R L K K E I S K F V D S S V T - b A K C K E D * R K R F Q N L L I L L * P - c R N E K K T K E R D F K I C C F F C N R -
GGAATTAACAACAAGAATATTAGCAACGAAAAAGAAGAAGAGGCTATCACAATCCTGATTC 1021+ 1080 CCTTAATTGTTGTTCTTATAATCGTTGCTTTTTCTTCTCGATAGTGTTAGGACTAAG
a G I N N K N I S N E K E E E L S Q S C F - b E L T T R I L A T K K K K S Y H N P D S - c N * Q Q E Y * Q R K R R R A I T I L I L -
TTAAAGATTTCAAAAATTCCAGGTAAGAGAGATACATTCATT
a L K I S K I P G K R D T F I K I H I L * - b * R F Q K F Q V R E I H S L K F I Y Y S - c K D F K N S R * E R Y I H * N S Y I I V -
TTTTTCATTTCACAGCTGTTATTTTCTTTATCTTAACAATATTTTTTTGATTAGCTGGAA 1141+ 1200 AAAAAGTAAAGTGTCGACAATAAAAGAAAATAGAATTGTTATAAAAAAACTAATCGACCTT
a F F I S Q L L F S F I L T I F F D * L E - b F S F H S C Y F L L S * Q Y F L I S W K - c F H F T A V I F F Y L N N I F C L A G S -
GTAAAAAGTATCAAATAAGAGAAGCGCTAGACTGAGGTAACTTAGCTTATTCACATTCAT 1201+ 1260 CATTTTCATAGTTTATTCTCTTCGCGATCTGACTCCATTGAATCGAATAAGTGTAAGTA
a V K S I K * E K R * T E V T * L I H I H - b * K V S N K R S A R L R * L S L F T F I - c K K Y Q I R E A L D C G N L A Y S H S * -
AGATCGACCTTCATATATCCAATACGATGATAAGGAAACAGCAGTCATCCGTTTTAAAAA 1261+ 1320 TCTAGCTGGAAGTATATAGGTTATGCTACTATTCCTTTGTCGTCAGTAGGCAAAATTTTT
a R S T F I Y P I R C * G N S S H P F * K - b D R P S Y I Q Y D D K E T A V I R F K N - c I D L H I S N T M I R K Q Q S S V L K I -
TAGTGCTATGAGGACTAAATTTTTAGAGTCAAGAAATGGAGCCGAAATCTTAATCAAAAA 1321+ 1380 ATCACGATACTCCTGATTTAAAAAATCTCAGTTCTTTACCTCGGCTTTAGAATTAGTTTTT
a * C Y E D * I F R V K K W S R N L N Q K -

FIG. 35C



```
GAATTGCGTCGATATTGCAAAAGAATCGAACTCTAAATCTTTCGTTAATAAGTATTACCA
1381 -----+ 1440
   {\tt CTTAACGCAGCTATAACGTTTTCTTAGCTTGAGATTTAGAAAGCAATTATTCATAATGGT}
  b
   I A S I L Q K N R T L N L S L I S I T N -
   {\tt ATCTTGATTGATGAGAGATTGACGAGGCAACTGCACAGAAGATCATTAAAGAAATAAA}
1441 -----+ 1500
   TAGAACTAACTTCTCTAACTGCTCCGTTGACGTGTCTTCTAGTAATTTCTTTATTT
  GTAACTTTTATTAATTAGAGAATAAACTAAATTACTAATATAGAGATCAGCGATCTTCAA
    ------+----+----+ 1560
   CATTGAAAATAATTAATCTCTTATTTGATTTAATGATTATCTCTAGTCGCTAGAAGTT
  V T F I N * R I N * I T N I E I S D L Q - * L L I R E * T K L L I * R S A I F N - N F Y * L E N K L N Y * Y R D Q R S S I -
b
   TTGACGAAATAAAAGCTGAACTAAAGTTAGACAATAAAAAATACAAACCTTGGTCAAAAT
1561 -----+ 1620
   b
  1621 -----+----+ 1680
   b
   GTACAGAAGTGAAGAATAAAAGATTTATTTTTTTCAATAATTTATTGAAAAGAGGGGTT
1681 -----+ 1740
   CATGTCTTCACTTCTTATTTTCTAAATAAAAAAAGTTATTAAATAACTTTTCTCCCCAA
  V Q K C R N K R F I F F N N L L K R G V
   R S E E I K D L F F S I I Y C K E G F -
T E V K K * K I Y F F Q * F I E K R G F -
  TTGGGGTTTTGGGG
1741 ------ 1762
  AACCCCAAAACCCCAAAACCCC
  L G F W G F G
W G F G V L G
G V L G F W
b
```

FIG. 35D



2	EVDVQNQADNHGIHSALKTCEEIKEAKTLYSWIQKVIRCRNQSQSHYKDL	51
19	::: :.:: : : .:: :: . ELELEMQENQNDIQVRVKIDDPKQYLVNVTAACLLQEGSYYQDK	62
52	EDIKIFAQTNIVATPRDYNEEDFKVIARKEVF.STGLMIELIDKCLVELL	100
63	:: .::::::::::::::::::::::::::::::::::	107
101	SSSDVSDRQKLQCFGFQLKGNQLAKTHLLTALSTQKQYFFQDEWNQVRAM:: ::: : . : .::	150
108		144
151	<pre>IGNELFRHLYTKYLIFQRTSEGTLVQFCGNNVFDHLKVNDKFDKKQKGGA :: : .: :.:: ::: ::</pre>	200
145	:: : : :::: :: :: :: :: :: :	181
201	ADMNEPRCCSTCKYNVKNEKDHFLNNINVPNWNNMKSRTRIFYCTHF	247
182	.:: .:: : . : :. .: SEFNEYQLGKYCTESQRKKTMFRYLSVTNKQKWDQTKKK	220
248	NRNNQFFKKHEFVSNKNNISAMDRAQTIFTNIFRFNRIRKKLKDKVIEKI	297
221	.: :::: .: : : :: . :. .RKENLLTKLQAIKESEDKSKRETGDIMNVEDAIKALKPAVMKKI	264
298	AYMLEKVKDFNFNYYLTKSCPLPENWRERKQKIENLINKTREEKSKYYEE	347
265	:	294
348	LFSYTTDNKCVTQFINEFFYNILPKDFLTGRNRKNFQKKVKKYVELNKHE	397
295	: : :: . LIKFCHISEPKERVYKILGKKYPKTEEEYKAAFGDSASAPFN.PE	338
398	LIHKNLLLEKINTREISWMQVETSAKHFYYFDHENIYVLWKLLRWIFEDL	447
339	: . : . : . : : . : : : LAGKRMKIEISKTWENELSAKGNTAEVWDNLISSNQLPYMAMLRNLSN	386
448	VVSLIRCFFYVTEQQKSYSKTYYYRKNIWDVIMKMSIADLKKETLAEVQE	497
387	: ::. ILKAGVSD	394
498	KEVEEWKKSLGFAPGKLRLIPKKTTFRPIMTFNKKIVNSDRKTTKLTTNT : .	547
395	······TTHS	398
548	KLLNSHLMLKTLKNRMFKDPFGFAVFNYDDVMKKYEEFVCKWKQVGQPKL	597
399	: . :. : IVINK	415
598	FFATMDIEKCYDSVNREKLSTFLKTTKLLSSDFWIMTAQILKRKNNIVID	647
416	:: :: :: :.: : :: FPLQFFSAIEAVN.EAVTKGFKAKKRENMNLKGQIEAVKEVVE	457
648	SKNFRKKEMKDYFRQKFQKIALEGGQYPTLFSVLENEQNDLNAKKTLIVE	697
458	: : . :: : :. KTDEEKKDMELEQTEEGEFVKVNEGIGKQYINSIELAIK	496
698	AKQRNYFKKDNLLQPVINICQYNYINFNGKFYKQTKGIPQGLCVSSILSS	747
497	::::::::::::::::::::::::::::::::::	546
748	FYYATLEESSLGFLRDESMNPENPNVNLLMRLTDDYLLITTQENNAVLFI : . . . : : : : : :	797
547	MVKQRCEKSSFYIFSSPSSQCNKCYLEVDL	576



	EKLINVSRENGFKFNMKK.LQTSFPLSPSKFAKYGMDSVEEQNIVQDYCD .::::: :::	
	PGDELRPSMQKLLQEKGKLGGGTDFPYECIDEWTKNKTHVD	
847	WIGISIDMKTLALMPNINLRIEGILCTLNLNMQTKKASMWLKKKLKSFLM . : :::: . .: NIVILSDMMIAEGYSDINVRGSSIVNSI	896
	NNITHYFRKTITTEDFANKTLNKLFISGGYKYMQCAKEYKD.HFKKNLAM .:.: ::: ::: PNIKIFAVDLEGYGKCLNLGDEFNENNYIKIFGM	
654	PNÍKIFAVDLEGYGKCLNLGDÉFNENNYIKIFGM	687
	SSMIDLEVSKIIYSVTRAFFKYLVCNIKDTIFGEEHYPDFFLSTLKHFIE . : :: :: SDSILKFISAKQGGANMVE	
688	SDSİLKFISAKQGGANMVÉ	706
	IFSTKKYIFNRVC 1008 :: .: :.::	
707	:: .: :.::. VIKNFALQKIG 717	

FIG. 36B

132	LSTQKQYFFQDEWNQVRAMIGNEL.FRHLYTKYLIFQRTSEGTLVQFC	178
1	: : : : :	43
179	GNNVFDHLKVNDKFDKKQKGGAADMNEPRCCSTCKYNVKNEKDHFLNNIN	228
44	:: :. : . .	84
229	VPNWNNMKSRTRIFYCTHFNRNNQFFKKHEFVSNKNNISAMDRAQTIFTN :: :: :. :	278
85	QIKQQVQLIKKVGSKVEKDLNLNEDENKKN	114
279	IFRFNRIRKKLKDKVIEKIAYMLEKVKDFNFNYYLTKSCPLPENWRERKQ :: ::::	328
115	GLSEQQVKEEQLRTITEEQVKYQNLVFNMDYQLDLNESGGHRRHRRETDY	164
329	KIENLINKTREEKSKYYEELFSYTTDNKCVTQFINE.FFYNILPKDFLTG	37
165	.::.:: .: .: : ::. DTEKWFEISHDQKNYVSIYANQKTSYCWWLKDYFNK	200
378	RNRKNFQKKVKKYVELNKHELIHKNLLLEKINTREISWMQVETSAKHFYY	42
201	NNYDHLNVSINRLETEAEFYAFDDFSQTIKLTNNSYQTVNID	242
428	FDHENIYVLWKLLRWIFEDLVVSLIRCFFYVTEQQKSYSKTYYYRKNI .: : :: : : : : .	475
243	VNFDNNLCILALLRFLLSLERFNILNIRSSYTRNQYNFEKIGELLETI	290
476	WDVIMKMSIADLKKETLAEVQEKEVEEWKKSLGFAPGKLRLIPKKTTFRP	525
291	:. : : :.:: . FAVVFSHRHLQGIHLQVPCEAFQYLVNSSSQISVKDSQLQ	330
526	<pre>IMTFNKKIVNSDRKTTKLTTNTKLLNSHLMLKTLKNRMFKDPFGFAVFNY : : . : </pre>	575
331	VYSFSTDLKLVD. TNKVQDYFKFLQEFPRLTHVSQQAIPVSATNAVENL	378

FIG. 37A

+



576	DDVMKKYEEFVCKWKQVGQPKLF FATMDIEKCYDS VNREK	615
379	: .: . . : .: : . :	426
516	LSTFLKTTKLLSSDFWIMTAQILKRKNNIVIDSKNFRKKEMK	657
427	LENLLLSIKQSKNLKFLRLNFYTYVAQETSRKQILKQATTIKNLKNNKNQ	476
558	DYFRQKFQKIALEGGQYPTLFSVLENEQNDLNAKKTLIVEAKQRNYFK	705
477	: : : :: . EETPETKDETPSESTSGMKFFDHLSELTELEDFSVNLQATQEIY	520
706	KDNLLQPVINICQYNYINFNGKFYKQTKGIPQGLCVSSILSSFYYATLEE	755
521	.DSLHKLLIRSTNLKKFKLSYKYEMEKSKMDTFIDLKNIYETLNN	564
756	SSLGFLRDESMNPENPNVNLLMRLTDDYLLITTQENNAVLFIEKLINVSR	305
565	: . : : .::: LKRCSVNISNPHGNISYELTNKDSTFYKFKLTLNQE	500
806	ENGFKFNMKKLQTSFPLSPSKFAKYGMDSVEEQNIVQDYCDWIGISIDMK	855
601	::: .:. .:. : .::: ::::LQHAKYTFKQNEFQFNNVKSAKIESSSLESLEDIDSLCKSIASCKNLQ	648
856	TLALMPNINLRIEGILCTLNLNMQTKKASMWLKKKLKSFLMNNITH	901
649	.:.: :: :. : ::: .: . . NVNIIASLLYPNNIQKNPFNKPNLLFFKQFEQLKNLENVSINC	691
902	YFRKTITTEDFANKTLNKLFISGGYKYMQCAKEYKDHFKKNLAMSSM .: . : .: : : ::	948
692	ILDQHILNSISEFLEKNKKIKAFILKRYYLLQYYLDYTKLFKTLQQLPEL	741
949	IDLEVSKIIYSVTRAFFKYLVCNIKDTIFGEEHY	982
742	:: : : :. . :: .: NQVYINQQLEELTVSEVHKQVWENHKQKAFYEPLCEFIKESSQTLQLIDF	791
983	PDFFLS TLKHFIEIFSTKKY IFNRVCMILKAKEAKLKSDQCQSLIQ .: : . :	1028
792	DQNTVSDDSIKKILESISESKYHHYLRLNPSQSSSLIKSENEEIQELLK	840

FIG. 37B

4	DIDLDDIENLLPNTFNKYSSSCSDKKGCKTLKSGSKSPSLTIPK	4
617	NVKSAKİĖSSSLESLEDIDSLČKSIASČKNLQNVNIIASLLYPNNIQKNP	66
48	LQKQLEFYFSDANLYNDSFLRKLVLKSGEQRVEIETLLM : : : ::: . . : : : : : :	8
667	FNKPNLLFFKOFEOLKNLENVSINCILDOHIINSISEFI.EKNKKİKAFII.	716



MEMDIDLDDIENLLPNTFNKYSSSCSDKKGCKTLKSGSKSPS : : :. : IELAIKIAVNKNLDEIKGHTAIFSDVSGSMSTSMSGGAKKYGSVRTCLEC	42 540
.LTIPKLQKQLEFYFSDANLYNDSFLRKLVLKSGEQRVEIETLL .: : : : : : ALVLGLMVKQRCEKSSFYIFSSPSSQCNKCYL.EVDLPGDELRPSMQKLL	85 589

FIG. 39

telomerase p43 LQKOLEFYFSDANLYNDSFIRKLVLKSGEQRVEIETLLM human La ICHOUEYYFGDFNLPRDKFIKEQI.KLDEGWVPLEIMIK Xenopus LaA ICEQIEYYFGDHNLPRDKFIKQQI.LLDDGWVPLETMIK Drosophila La ILROVEYYFGDANLNRDKFIREQIGKNEDGWVPLSVLVT S. c. Lhplp CLKQVEFYFSEFNFPYDRFIRTTAEK.NDGWVPISTIAT

FIG. 41

1 aactcattta attactaatt taatcaacaa gattgataaa aagcagtaaa taaaacccaa 61 tagatttaat ttagaaagta tcaattgaaa aatggaaatt gaaaacaact aagcacaata 121 gccaaaagcc gaaaaattgt ggtgggaact tgaattagag atgcaagaaa accaaaatga 181 tatataagtt agggttaaga ttgacgatcc taagcaatat ctcgtgaacg tcactgcagc 241 atgtttgttg taggaaggta gttactacta agataaagat gaaagaagat atatcatcac 301 taaagcactt cttgaggtgg ctgagtctga tcctgagttc atctgctagt tggcagtcta 361 catcogtaat gaactttaca toagaactac cactaactac attgtagcat titgtgttgt 421 ccacaagaat actcaaccat tcatcgaaaa gtacttcaac aaagcagtac ttttgcctaa 481 tgacttactg gaagtctgtg aatttgcata ggttctctat atttttgatg caactgaatt 541 caaaaatttg tatcttgata ggatactttc ataagatatt cgtaaggaac tcactttccg 601 taagtgttta caaagatgcg tcagaagcaa gttttctgaa ttcaacgaat actaacttgg 661 taagtattgc actgaatcct aacgtaagaa aacaatgttc cgttacctct caqttaccaa 721 caagtaaaag tgggattaaa ctaagaagaa gagaaaagag aatctcttaa ccaaacttta 781 ggcaataaag gaatctgaag ataagtccaa gagagaaact ggagacataa tgaacgttga 841 agatgcaatc aaggctttaa aaccagcagt tatgaagaaa atagccaaga gatagaatgc 901 catgaagaaa cacatgaagg cacctaaaat tectaaetet accttggaat caaagtaett 961 gacetteaag gateteatta agttetgeea tatttetgag eetaaagaaa gagtetataa 1021 gateettggt aaaaaatace etaagaeega agaggaatae aaageageet ttggtgatte 1081 tgcatctgca cccttcaatc ctgaattggc tggaaagcgt atgaagattg aaatctctaa 1141 aacatgggaa aatgaactca gtgcaaaagg caacactgct gaggtttggg ataatttaat 1201 ttcaagcaat taactcccat atatggccat gttacgtaac ttgtctaaca tcttaaaagc 1261 cggtgittca gatactacac actelattgt gatcaacaag atttgtgagc ccaaggccgt 1321 tgagaactcc aagatgttcc ctcttcaatt ctttagtgcc attgaagctg ttaatgaagc 1381 agttactaag ggattcaagg ccaagaagag agaaaatatg aatcttaaag gtcaaatcga 1441 agcagtaaag gaagttgttg aaaaaaccga tgaagagaag aaagatatgg agttggagta 1501 aaccgaagaa ggagaatttg ttaaagtcaa cgaaggaatt ggcaagcaat acattaactc 1561 cattgaactt gcaatcaaga tagcagttaa caagaattta gatgaaatca aaggacacac 1621 tgcaatcttc tctgatgttt ctggttctat gagtacctca atgtcaggtg gagccaagaa 1681 gtatggttcc gttcgtactt gtctcgagtg tgcattagtc cttggtttga tggtaaaata 1741 acgttgtgaa aagtcctcat tctacatctt cagttcacct agttctcaat gcaataagtg 1801 ttacttagaa gttgatetee etggagaega acteegteet tetatgtaaa aaettttgea 1861 agagaaagga aaacttggtg gtggtactga tttcccctat gagtgcattg atgaatggac 1921 aaagaataaa actcacgtag acaatatcgt tattttgtct gatatgatga ttgcagaagg 1981 atatteagat ateaatgtta gaggeagtte cattgttaac ageateaaaa agtacaagga 2041 tgaagtaaat eetaacatta aaatetttge agttgaetta gaaggttaeg gaaagtgeet 2101 taatetaggt gatgagttea atgaaaacaa etacateaag atatteggta tgagegatte 2161 aatcttaaag ttcatttcag ccaagcaagg aggagcaaat atggtcgaag ttatcaaaaa 2221 ctttgccctt caaaaaatag gacaaaagtg agtttcttga gattcttcta taacaaaaat 2281 ctcacccac ttttttgttt tattgcatag ccattatgaa atttaaatta ttatctatt 2341 atttaagtta cttacatagt ttatgtatcg cagtctatta gcctattcaa atgattctqc 2401 aaagaacaaa aaagattaaa a



Motif A Motif

7-GIRYQYNVLPQGWKGSPAIFQSSMTKILEPFRKQN GQPKLFFATMDIEKCYDSVNREKLSTFLKTTKLL-100-KFYKQTKGIP**QG**LCV**S**SILSSFYYATLEESSLGFL KNRNLHCTYDDYKKAFDSIPHSWLIQVLEIYKIN- 28-RQIAIKKGIY**QG**DSLSPLWFCLALNPLSHQLHNDR FGGSNWFREV**D**LKKCFDTISHDLIIKELKRYISD- 26-HVPVGPRVCV**QG**APT**S**PALCNAVLLRLDRRLAGLA VLPELYFMKFDVKSCYDSIPRMECMRILKDALKN- 68-KCYIREDGLFQGSSLSAPIVDLVYDDLLEFYSEFK h-hLGh-h -16-HLIYMDDIKLYAKNDKE-0-MKKLIDTTTIFSNDISMQFGLDKCKT-25-KCLYKYLGFQQ 4-IYQYMDDLYVGSHLEIG-1-HRTKIEELRQHLLRWGLTTPDKKHQK- 0-EPPFLWMGYEL
 8-ILKLADDFLIISTDQQQ........VINIKKLAMGGFQKYNARANR-41-IRSKSSKGIFR -14-LMRLTDDYLLITTQENN-0-AVLFIEKLINVSRENGFKFNMKKLQT-23-QDYCDWIGISI -55-YVRYADDILIGVLGSKN-2-KIIKRDLNNFLNS.LGLTINEEKTLI- 4-ETPARFLGYÑĨ Motif E h---+-QG----RP Gh-h---K Motif D LKKKKSVTVL**D**VGDAYFSVPLDEDFRKYTAFTIPh--hDh---h h--YhDDhhh Motif C al S.c. (groupII) Dong (LINE) al S.c. (groupII) telomerase p123 telomerase p123 Dong (LINE) Consensus Consensus L8543.12 L8543.12 HIV-RT HIV-RT

7.0.7

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MEIENNQAQQPKAEKLWWELELEMQENQNDIQVRVKIDDPKQYL VNVTAACLLQEGSYYQDKDERRYIITKALLEVAESDPEFICQLA VYIRNELYIRTTTNYIVAFCVVHKNTQPFIEKYFNKAVLLPNDL LEVCEFAOVLYIFDATEFKNLYLDRILSODIRKELTFRKCLORC VRSKFSEFNEYOLGKYCTESORKKTMFRYLSVTNKOKWDOTKKK RKENLLTKLQAIKESEDKSKRETGDIMNVEDAIKALKPAVMKKI AKRQNAMKKHMKAPKIPNSTLESKYLTFKDLIKFCHISEPKERV YKILGKKYPKTEEEYKAAFGDSASAPFNPELAGKRMKIEISKTW ENELSAKGNTAEVWDNLISSNQLPYMAMLRNLSNILKAGVSDTT HSIVINKICEPKAVENSKMFPLQFFSAIEAVNEAVTKGFKAKKR ENMNLKGQIEAVKEVVEKTDEEKKDMELEOTEEGEFVKVNEGIG KQYINSIELAIKIAVNKNLDEIKGHTAIFSDVSGSMSTSMSGGA KKYGSVRTCLECALVLGLMVKQRCEKSSFYIFSSPSSQCNKCYL EVDLPGDELRPSMQKLLQEKGKLGGGTDFPYECIDEWTKNKTHV DNIVILSDMMIAEGYSDINVRGSSIVNSIKKYKDEVNPNIKIFA VDLEGYGKCLNLGDEFNENNYIKIFGMSDSILKFISAKQGGANM VEVIKNFALQKIGQK

FIG. 43

MSRRNQKKPQAPIGNETNLDFVLQNLEVYKSQIEHYKTQQQQIK EEDLKLLKFKNODODGNSGNDDDDEENNSNKOOELLRRVNOIKO QVQLIKKVGSKVEKDLNLNEDENKKNGLSEQQVKEEQLRTITEE QVKYQNLVFNMDYQLDLNESGGHRRHRRETDYDTEKWFEISHDQ KNYVSIYANQKTSYCWWLKDYFNKNNYDHLNVSINRLETEAEFY AFDDFSQTIKLTNNSYQTVNIDVNFDNNLCILALLRFLLSLERF NILNIRSSYTRNQYNFEKIGELLETIFAVVFSHRHLQGIHLQVP CEAFQYLVNSSSQISVKDSQLQVYSFSTDLKLVDTNKVQDYFKF LQEFPRLTHVSQQAIPVSATNAVENLNVLLKKVKHANLNLVSIP TQFNFDFYFVNLQHLKLEFGLEPNILTKQKLENLLLSIKQSKNL KFLRLNFYTYVAQETSRKQILKQATTIKNLKNNKNQEETPETKD ETPSESTSGMKFFDHLSELTELEDFSVNLQATQEIYDSLHKLLI RSTNLKKFKLSYKYEMEKSKMDTFIDLKNIYETLNNLKRCSVNI SNPHGNISYELTNKDSTFYKFKLTLNQELQHAKYTFKQNEFQFN NVKSAKIESSSLESLEDIDSLCKSIASCKNLQNVNIIASLLYPN NIQKNPFNKPNLLFFKQFEQLKNLENVSINCILDOHILNSISEF LEKNKKIKAFILKRYYLLQYYLDYTKLFKTLQQLPELNQVYINQ QLEELTVSEVHKQVWENHKQKAFYEPLCEFIKESSQTLQLIDFD QNTVSDDSIKKILESISESKYHHYLRLNPSOSSSLIKSENEEIO ELLKACDEKGVLVKAYYKFPLCLPTGTYYDYNSDRW

FIG. 45

MKILFEFIQDKLDIDLQTNSTYKENLKCGHFNGLDEILTTCFAL PNSRKIALPCLPGDLSHKAVIDHCIIYLLTGELYNNVLTFGYKI ARNEDVNNSLFCHSANVNVTLLKGAAWKMFHSLVGTYAFVDLLI NYTVIQFNGQFFTQIVGNRCNEPHLPPKWVQRSSSSSATAAQIK QLTEPVTNKQFLHKLNINSSSFFPYSKILPSSSSIKKLTDLREA IFPTNLVKIPQRLKVRINLTLQKLLKRHKRLNYVSILNSICPPL EGTVLDLSHLSRQSPKERVLKFIIVILOKLLPOEMFGSKKNKGK IIKNLNLLLSLPLNGYLPFDSLLKKLRLKDFRWLFISDIWFTKH NFENLNQLAICFISWLFRQLIPKIIQTFFYCTEISSTVTIVYFR HDTWNKLITPFIVEYFKTYLVENNVCRNHNSYTLSNFNHSKMRI IPKKSNNEFRIIAIPCRGADEEEFTIYKENHKNAIQPTQKILEY LRNKRPTSFTKIYSPTQIADRIKEFKQRLLKKFNNVLPELYFMK FDVKSCYDSIPRMECMRILKDALKNENGFFVRSQYFFNTNTGVL KLFNVVNASRVPKPYELYIDNVRTVHLSNQDVINVVEMEIFKTA $\verb|LWVEDKCYIREDGLFQGSSLSAPIVDLVYDDLLEFYSEFKASPS|$ QDTLILKLADDFLIISTDQQQVINIKKLAMGGFQKYNAKANRDK ILAVSSQSDDDTVIQFCAMHIFVKELEVWKHSSTMNNFHIRSKS SKGIFRSLIALFNTRISYKTIDTNLNSTNTVLMQIDHVVKNISE CYKSAFKDLSINVTQNMQFHSFLQRIIEMTVSGCPITKCDPLIE YEVRFTILNGFLESLSSNTSKFKDNIILLRKEIQHLQAYIYIYI HTVN

FIG. 46

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1 tcaatactat taattaataa ataaaaaaaa gcaaactaca aagaaaatgt caaggcgtaa
 61 ctaaaaaaag ccataggctc ctataggcaa tgaaacaaat cttgattttg tattacaaaa 121 tctagaagtt tacaaaagcc agattgagca ttataagacc tagtagtaat agatcaaaga
 181 ggaggatete aagettttaa agtteaaaaa ttaagattag gatggaaact etggeaacga
 241 tgatgatgat gaagaaaaca actcaaataa ataataagaa ttattaagga gagtcaatta
 301 gattaagtag caagtttaat tgataaaaaa agttggttct aaggtagaga aagatttqaa
 361 tttgaacgaa gatgaaaaca aaaagaatgg actttctgaa tagcaagtga aagaagagta
 421 attaagaacg attactgaag aataggttaa gtattaaaat ttagtattta acatggacta
 481 ccagttagat ttaaatgaga gtggtggcca tagaagacac agaagagaaa cagattatga
 541 tactgaaaaa tggtttgaaa tatctcatga ccaaaaaaat tatgtatcaa tttacgccaa
 601 ctaaaagaca tcatattgtt ggtggcttaa agattatttt aataaaaaca attatgatca
 661 tettaatgta ageattaaca gaetagaaac tgaageegaa ttetatgeet ttgatgattt
 721 ttcacaaaca atcaaactta ctaataattc ttactagact gttaacatag acgttaattt
 781 tgataataat ctctgtatac tcgcattgct tagattitta ttatcactag aaagattcaa
 841 tattttgaat ataagatett ettataeaag aaattaatat aattttgaga aaattggtga
901 getaettgaa actatetteg eagttgtett tteteatege eacttaeaag geatteattt
961 acaagtteet tgegaagegt tetaatattt agttaaetee teateataa ttagegttaa
1021 agatagetaa ttataggtat actetttete tacagaetta aaattagttg acaetaacaa
1081 agtocaagat tattttaagt tottataaga attooctogt ttgactoatg taagotagta
1141 ggctatccca gttagtgcta ctaacgctgt agagaacctc aatgttttac ttaaaaaqqt
1201 caagcatgct aatcttaatt tagtttctat ccctacctaa ttcaattttg atttctactt
1261 tgttaattta taacatttga aattagagtt tggattagaa ccaaatattt tgacaaaaca
1321 aaagettgaa aatetaettt tgagtataaa ataateaaaa aatettaaat tittaagatt
1381 aaacttttac acctacgttg cttaagaaac ctccagaaaa cagatattaa aacaagctac
1441 aacaatcaaa aatctcaaaa acaataaaaa tcaagaagaa actcctgaaa ctaaagatga
1501 aactccaagc gaaagcacaa gtggtatgaa attttttgat catctttctg aattaaccga
1561 gcttgaagat ttcagcgtta acttgtaagc tacccaagaa atttatgata gcttgcacaa
1621 acttitgatt agatcaacaa atttaaagaa gttcaaatta agttacaaat atgaaatgga
1681 aaagagtaaa atggatacat tcatagatct taagaatatt tatgaaacct taaacaatct
1741 taaaagatgc tctgttaata tatcaaatcc tcatggaaac atttcttatg aactgacaaa 1801 taaagattct acttttata aatttaagct gaccttaaac taagaattat aacacgctaa
1861 gtatactttt aagtagaacg aattttaatt taataacgtt aaaagtgcaa aaattgaatc
1921 ttcctcatta gaaagcttag aagatattga tagtctttgc aaatctattg cttcttgtaa
1981 aaatttacaa aatgttaata ttatcgccag tttgctctat cccaacaata tttagaaaaa
2041 teettteaat aageeeaate ttetatttt caageaattt gaataattga aaaatttgga
2101 aaatgtatet ateaaetgta tiettgatea geatataett aattetatit eagaattett
2161 agaaaagaat aaaaaaataa aagcattcat tttgaaaaga tattatttat tacaatatta
2221 tottgattat actaaattat ttaaaacact tcaatagtta cotgaattaa attaagttta
2281 cattaattag caattagaag aattgactgt gagtgaagta cataagtaag tatgggaaaa
2341 ccacaagcaa aaagctttct atgaaccatt atgtgagttt atcaaagaat catcctaaac
2401 cctttagcta atagattttg accaaaacac tgtaagtgat gactctatta aaaagatttt
2461 agaatctata totgagtota agtatoatoa ttatttgaga ttqaaccota qttaatctaq
2521 cagtttaatt aaatctgaaa acgaagaaat ttaagaactt ctcaaagctt gcgacgaaaa
2701 tgaatatttc tttgcttatt atttgaataa tacatacaat agtcattttt agtgttttga
2761 atatatttta gttatttaat tcattatttt aagtaaataa ttattttca atcattttt
2821 aaaaaatcq
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FIG. 44





Oxytricha Euplotes LCVSYILSSFYYANLEENALQFLRKESMDPEKPETNLLMRLT LCVSSILSSFYYATLEESSLGFLRDESMNPENPNVNLLMRLT

FIG. 47

ATTTATACTCATGAAAATCTTATTCGAGTTCATTCAAGACAAGCTTGACATTGATCTACA GACCAACAGTACTTACAAAGAAAATTTAAAATGTGGTCACTTCAATGGCCTCGATGAAAT TCTAACTACGTGTTTCGCACTACCAAATTCAAGAAAAATAGCATTACCATGCCTTCCTGG TGACTTAAGCCACAAAGCAGTCATTGATCACTGCATCATTTACCTGTTGACGGGCGAATT ATACAACAACGTACTAACATTTGGCTATAAAATAGCTAGAAATGAAGATGTCAACAATAG TCTTTTTTGCCATTCTGCAAATGTTAACGTTACGTTACTGAAAGGCGCTGCTTGGAAAAT GTTCCACAGTTTGGTCGGTACATACGCATTCGTTGATTATTGATCAATTATACAGTAAT TCAATTTAATGGGCAGTTTTTCACTCAAATCGTGGGTAACAGATGTAACGAACCTCATCT GCCGCCCAAATGGGTCCAACGATCATCCTCATCATCCGCAACTGCTGCGCAAATCAAACA ACTTACAGAACCAGTGACAAATAAACAATTCTTACACAAGCTCAATATAAATTCCTCTTC TTTTTTTCCTTATAGCAAGATCCTTCCTTCATCATCATCTATCAAAAAGCTAACTGACTT GAGAGAAGCTATTTTCCCACAAATTTGGTTAAAATTCCTCAGAGACTAAAGGTACGAAT TAATTTGACGCTGCAAAAGCTATTAAAGAGACATAAGCGTTTGAATTACGTTTCTATTTT GAATAGTATTTGCCCACCATTGGAAGGGACCGTATTGGACTTGTCGCATTTGAGTAGGCA ATCACCAAAGGAACGAGTCTTGAAATTTATCATTGTTATTTTACAGAAGTTATTACCCCA AGAAATGTTTGGCTCAAAGAAAAATAAAGGAAAAATTATCAAGAATCTAAATCTTTTATT AAGTTTACCCTTAAATGGCTATTTACCATTTGATAGTTTGTTGAAAAAGTTAAGATTAAA GGATTTTCGGTGGTTGTTCATTTCTGATATTTGGTTCACCAAGCACAATTTTGAAAACTT GAATCAATTGGCGATTTGTTTCATTTCCTGGCTATTTAGACAACTAATTCCCAAAATTAT ACAGACTTTTTTTTACTGCACCGAAATATCTTCTACAGTGACAATTGTTTACTTTAGACA TGATACTTGGAATAAACTTATCACCCCTTTTATCGTAGAATATTTTAAGACGTACTTAGT CGAAAACAACGTATGTAGAAACCATAATAGTTACACGTTGTCCAATTTCAATCATAGCAA AATGAGGATTATACCAAAAAAAGTAATAATGAGTTCAGGATTATTGCCATCCCATGCAG AGGGGCAGACGAAGAATTCACAATTTATAAGGAGAATCACAAAAATGCTATCCAGCC CACTCAAAAAATTTTAGAATACCTAAGAAACAAAAGGCCGACTAGTTTTACTAAAATATA TTCTCCAACGCAAATAGCTGACCGTATCAAAGAATTTAAGCAGAGACTTTTAAAGAAATT TAATAATGTCTTACCAGAGCTTTATTTCATGAAATTTGATGTCAAATCTTGCTATGATTC CATACCAAGGATGGAATGTATGAGGATACTCAAGGATGCGCTAAAAAATGAAAATGGGTT TTTCGTTAGATCTCAATATTTCTTCAATACCAATACAGGTGTATTGAAGTTATTTAATGT TGTTAACGCTAGCAGAGTACCAAAACCTTATGAGCTATACATAGATAATGTGAGGACGGT TCATTTATCAAATCAGGATGTTATAAACGTTGTAGAGATGGAAATATTTAAAACAGCTTT GTGGGTTGAAGATAAGTGCTACATTAGAGAAGATGGTCTTTTTCAGGGCTCTAGTTTATC TGCTCCGATCGTTGATTTGGTGTATGACGATCTTCTGGAGTTTTATAGCGAGTTTAAAGC CAGTCCTAGCCAGGACACATTAATTTTAAAACTGGCTGACGATTTCCTTATAATATCAAC AGACCAACAGCAAGTGATCAATATCAAAAAGCTTGCCATGGGCGGATTTCAAAAATATAA TGCGAAAGCCAATAGAGACAAAATTTTAGCCGTAAGCTCCCAATCAGATGATGATACGGT TATTCAATTTTGTGCAATGCACATATTTGTTAAAGAATTGGAAGTTTGGAAACATTCAAG CACAATGAATAATTTCCATATCCGTTCGAAATCTAGTAAAGGGATATTTCGAAGTTTAAT AGCGCTGTTTAACACTAGAATCTCTTATAAAACAATTGACACAAATTTAAATTCAACAAA CACCGTTCTCATGCAAATTGATCATGTTGTAAAGAACATTTCGGAATGTTATAAATCTGC TTTTAAGGATCTATCAATTAATGTTACGCAAAATATGCAATTTCATTCGTTCTTACAACG CATCATTGAAATGACAGTCAGCGGTTGTCCAATTACGAAATGTGATCCTTTAATCGAGTA TGAGGTACGATTCACCATATTGAATGGATTTTTGGAAAGCCTATCTTCAAACACATCAAA ATTTAAAGATAATATCATTCTTTTGAGAAAGGAAATTCAACACTTGCAAGC



AKFLHWLMSVYVVELLRSFFYVTETTFQKNRLFFYRKSVWSKLQSIGIRQHLKR VQLRDVSEAEVRQHREARPALLTSRLRFIPKPDGLRPIVNMDYVVGARTFRREK RAERLTSRVKALFSVLNYERA

FIG. 49

GCCAAGTTCCTGCACTGGCTGATGAGTGTGTACGTCGTCGAGCTGCTCAGGTC
TTTCTTTTATGTCACGGAGACCACGTTTCAAAAGAACAGGCTCTTTTTTCTACC
GGAAGAGTGTCTGGAGCAAGTTGCAAAGCATTGGAATCAGACAGCACTTGAAG
AGGGTGCAGCTGCGGGACGTGTCGGAAGCAGAGGTCAGCAGCATCGGGAAGC
CAGGCCCGCCTGACGGTCCAGACTCCGCTTCATCCCCAAGCCTGACGGC
TGCGGCCGATTGTAACATGGACTACGTCGTGGGAGCCAGAACGTTCCGCAGA
GAAAAGAGGGCCGAGCGTCTCACCTCGAGGGTGAAGGCACTGTTCAGCGTGCT
CAACTACGAGCGGCGCG

FIG. 50

MTEHHTPKSRILRFLENQYVYLCTLNDYVQLVLRGSPASSYSNICERLRSDVQTSFSIFLHSTVVGF
DSKPDEGVQFSSPKCSQSELIANVVKQMFDESFERRNLLMKGFSMNHEDFRAMHVNGVQNDLVSTF
PNYLISILESKNWQLLLEIIGSDAMHYLLSKGSIFEALPNDNYLQISGIPLFKNNVFEETVSKKRKR
TIETSITQNKSARKEVSWNSISISRFSIFYRSSYKKFKQDLYFNLHSICDRNTVHMWLQWIFPRQFG
LINAFQVKQLHKVIPLVSQSTVVPKRLLKVYPLIEQTAKRLHRISLSKVYNHYCPYIDTHDDEKILS
YSLKPNQVFAFLRSILVRVFPKLIWGNQRIFEIILKDLETFLKLSRYESFSLHYLMSNIKISEIEWL
VLGKRSNAKMCLSDFEKRKQIFAEFIYWLYNSFIIPILQSFFYITESSDLRNRTVYFRKDIWKLLCR
PFITSMKMEAFEKINENNVRMDTQKTTLPPAVIRLLPKKNTFRLITNLRKRFLIKMGSNKKMLVSTN
QTLRPVASILKHLINEESSGIPFNLEVYMKLLTFKKDLLKHRMFGRKKYFVRIDIKSCYDRIKQDLM
FRIVKKKLKDPEFVIRKYATIHATSDRATKNFVSEAFSYFDMVPFEKVVQLLSMKTSDTLFVDFVDY
WTKSSSEIFKMLKEHLSGHIVKIGNSQYLQKVGIPQGSILSSFLCHFYMEDLIDEYLSFTKKKGSVL
LRVVDDFLFITVNKKDAKKFLNLSLRGFEKHNFSTSLEKTVINFENSNGIINNTFFNESKKRMPFFG
FSVNMRSLDTLLACPKIDEALFNSTSVELTKHMGKSFFYKILRSSLASFAQVFIDITHNSKFNSCCN
IYRLGYSMCMRAQAYLKRMKDIFIPQRMFITDLLNVIGRKIWKKLAEILGYTSRRFLSSAEVKWLFC
LGMRDGLKPSFKYHPCFEQLIYQFQSLTDLIKPLRPVLRQVLFLHRRIAD



actcaataacaataccaagtcaaattccaatatgaaggtgttattagtgatcgataatatttctattttatcggtcgtta ccaagtataaggacaaaaagaacaacttccttcccctaaagacttttactttaatttaattttcaaatatttc ttaacatggagccttacactttagatgagtcacgtcgcatgatgagtattttggtattcatccaccacgtttgccttgaaaag gttgataattatttgcaaaatcatgtccttagtggtggtaatccgcgaaagttttttgatgcttgcacacgtctagcatg ataatctaaattagtttcgcttataattgatagtagtagaaagattggtgattctactcgtgtaatgttattagtttaăa CAGTGATGCCATGCCATTACTTATTATCCAAAGGAAGTATTTTTGÄGGCTCTTCCAAATGACAATTACCTTCAGATTTCTG ACAAACAGCAAAGCGACTCCATCGTATTTCTCTATCAAAAGTTTACAACCATTATTGCCCATATATTGACACCCATG ggtaccgatttactttcctttcttcataagctaattgcttcctcgaacgctcctaaatctctggaaatattttacaaga agctcttggagtagctcacagaaatccttacaaatcttctgatgagactatattagattcattacagtccgtgcatattc attgagatattcaaaaaatttctatcactacaactcctttaacgcggttttatttttctatttctattctcattctcatdttgt ccaaatatgtatcatctcgtattaggctttttcccgttttactcctggaatcgtacctttttcactattccccctaatga attctaattgtgaaatatttacctgcaattactgtttcaaagagattgtatttaaccgataaagAATCATGAAGATTTTC GAGCCATGCATGTAAACGGAGTACAAAATGATCTCGTTTCTACTTTTCCTAATTACCTTATATCTATACTTGAGTCAAAA AATTGGCAACTTTTGTTAGAAATgtaaataccggttaagatgttgcgcactttgaacaagactgacaagtatagTATCGG CAAAATAAAAGCGCCCGCAAAGAAGTTTCCTGGAATAGCATTTCAATTAGTAGGTTTAGCATTTTTACAGGTTCATC caatgtactttacttctaatctattattagcagATGGGTTCAAACAAAAAAAATGTTAGTCAGTACGAACCAAACTTTACG GACCGAACCATACCCCCAAAAGCAGGATTCTTCGCTTTCTAGAGAATCAATATGTATACCTATGTACCTTAAATGATT ATGTACAACTTGTTTTGAGAGGGTCGCCGGCAAGCTCGTATAGCAATATATGCGAACGCTTGAGAAGCGATGTACAAACG TCCTTTTCTATTTTTCTTCATTCGACTGTAGTCGGCTTCGACAGTAAGCCAGATGAAGGTGATGAAGTTCAATTTTCTTCTCCAAA ATGCTCACAGTCAGAGgtatatatttttttttttttttttttttttttctattcgggatagctaatatgggcagCTAATAGC GTGATCGGAACACAGTACACATGTGGCTTCAATGGATTTTTCCAAGGCAATTTGGACTTATAAACGCATTTCAAGTGAAG CAATTGCACAAAGTGATTCCACTGGTATCACAGAGTACAGTTGTGCCCCAAACGTCTCCTAAAGGTATACCCTTTAATTGA **ATGAAAAATCCTTAGTTATTCCTTAAAGCCGAACCAGGTGTTTGCGTTTCTTCGATCCATTCTTGTTCGAGTGTTTCCT** accaqACCTCGAAACTTTCTTGAAATTATCGAGATACGAGTCTTTTTAGTTTACATTATTTAATGAGTAACATAAGGtaa tatgccaaatttttttaccattaattaacaatcaqATTTCAGAAATTGAATGGCTAGTCCTTGGAAAAAGGTCAAATGCG **ACCTGTGGCATCGATACTGAAACATTTAATCAATGAAGAAAGTAGTGGTATTCCATTTAACTTGGAGGTTTACATGAAGC** ggttcgcttacttttaatcgtggtactgttttagctgctacttctagccaaccgcgtgtttctaccccgtcattggatat GAATGTTGTAAAACAGATGTTCGATGAAAGTTTTGAGCGTCGAAGGAATCTACTGATGAAAGGGTTTTTCCATGGtaaggt TAAGAAGTTTAAGCAAGgtaactaatactgttatccttcataactaattttagATCTATATTTAACTTACACTCTATTT AAATTAATCTGGGGTAACCAAAGGATATTTGAGATAATATTAAAAGgtattgtataaaatttattaccactaacgatttt aaaatgtgcttaagtgattttgagaaacgcaagcaaatatttgcggaattcatctagctgtgtacaattcgtttataat ACCTATTTACAATCTTTTTTTTTATATCACTGAATCAAGTGATTTACGAAATCGAACTGTTTATTTTAGAAAAGATATT tttgcaaaaagctaatattttcagAACAATGTTAGGATGGATACTCAGAAAACTACTTTGCCTCCAGCAGTTATTCGTC TATTÄCCTAAGÄAGAATACCTTTCĞTCTCATTACGAATTTAAGAAAAAGATTCTTAATAAAGGtattaaatttttggtcat



AAACTTTGTTAGTGAGGCGTTTTCCTATTGtaagtttattttttcattggaattttttaacaaattctttttagTTGAT ATGGTGCCTTTTGAAAAGTCGTGCAGTTACTTTCTATGAAAACATCAGÄTACTTTGTTGTTGTTTTTTTGTGGATTATTG GACCAAAAGTTCTTCTGAAATTTTTAAAATGCTCAAGGAACATCTCTCTGGACACTTGTTAAGGtataccaattgttga AGCTGACGAAACATATGGGGAAATCTTTTTTTACAAATTCTAAGgtatactgtaactgaataatagctgacaaata atcaqATCGAGCCTTGCATCCTTTGCACAAGTATTTATTGACATTACCCACAATTCAAAATTCAAATTCTTGCTGCAATAT TGTTCATAACGGgtgagtacttatttaactagaaaagtcattaattaaccttagATCTTTGAATGTTATTGGAAGAAA TATCATCCATGCTTCGAACAGCTAATATACCAATTTCAGTCATTGACTGATCTTATCAAGCCGGCTAAGACCAGTTTTGCG ACAGGTGTTATTTTTACATAGAAGAATAGCTGATTAAtgtcattttcaattttattatatacatcctttattactggtgtc agtitigaatactaatagctcattiaatgtcitatataaaggittigttitttccigacticaatittgcatgggtgaaaag TTCTTACTTTTAAGAAGGATCTTCTTAAGCACCGAATGTTTGGgtaattatataatgcgcgattcctcattattaatttt gcagGCGTAAGAAGTATTTTGTACGGATAGATATAAAATCCTGTTATGATCGAATAAAGCAAGATTTGATGTTTCGATT ĞTTÄAAAAGAAACTCAAGGATCCCGAATTTGTAATTCGAAAGTATGCAACCATACATGCAACAAGTGACCGAGCTACAAA attgtaataacactaatgaaactaqATAGGAAATTCTCAATACCTTCAAAAAGTTGGTATCCTTCAGGGGCTCAATTCTGT TTACGAGTAGTCGACGATTTCCTCTTTATAACAGTTAATAAAAGGATGCAAAAAATTTTTGAATTTTATCTTTAAGAGG tgagttgctgtcattcctaagttctaaccgttgaagGATTTGAGAAACACAATTTTTCTACGAGCCTGGAGAAACAGTĀ CTCTGTGAACATGAGGTCTCTTGATACATTGTTAGCATGTCCTAAAATTGATGAAGCCTTATTTAACTCTACATCTGTAG <u>AATTTGGAAAAAĞTTGĞCCGAAATATTAGGATÄTACGÄGTAGGCGTTTCTTGTCČTCTGCAGAAGTCAAATGGtacgtgt</u> cggtctcgagacttcagcaatattgacacatcagGCTTTTTGTCTTGGAATGAGAGATGGTTTGAAACCCTCTTTCAAA cccattaaacgggagtggttaaaacattaaaagtaatacatgaggctaatctcctttcatttagaataaggaaagtggttt tctataatgaataatgcccgcactaatgcaaaaagacgaagattatcttctaaaacaagggggattaagcatatccgaagg aaaagagagtaatatacccagtgttgttgaagaaagcaaggataatttggaacaagcttctgcagatgacaggctaaatt aaatagtgttaagccattattggattccgaaatagccaaatttcttggttcctcaaagcggaagtctaaaagaacttattg gctgaggagaagcctaattttttgcaaaaaagaaatatcattgggagacatctcttgatgaatcagatgcggagagtat stccagcggatccttgatgtcaataacttctatttctgaaatgtatggtcctactgtcgcttcgacttctcgtagctctc aagottatgaggottcaaaaaotcotcotgatttaaaggaggaatottccaccgatgaggaaatggatagcttatcagot ATAAACTTTGAAAATAGTAATGGGATAATAAAAAAATACTTTTTTAATGAAAGCAAGAAAAGAATGCCATTCTTCGGTT cgcagttaagtgaccaaaggtacc

FIG. 52E



WNKLIT PFIVE YFK-TYLVEN 40 IWDVI-MKMS IADLKK ETLAEVQE 43 IWKLVSKLTI VKVRIQFSEK NKQMKNNFYQ 44	KLV 50	HSKM RIIPKKSNNE FRLIBIPCRG 79 PGKG RLIPKKTT FRPIMTFNKK 78 QGKL RIIPKKGS FRPIMTFLRK 92	FGKL RIIPKK FRPIMTF.RK 100	QKILEYLRNK RPTSFTKIYS PTQIADRIKE 129 HLMLKTLKNRMFK -DPFGFAVFN 120 QLVFRNLKDML-G -QKIGYSVFD 130	LKN 150.VF. 150	MKFD VKSCYD 157 ATMD IEKCYD 155 VTL 158
FFYCTEISST VTIVYFRHDT WNKLIT PFIVE YFK-TYLVEN FFYVTEQQKS YSKTYYYRKN IWDVI-MKMS IADLKK ETLAEVQE KHKE GSQIFYYRKP IWKLVSKLTI VKVRIQFSEK NKQMKNNFYQ	FFY.TEKSYYYRK. IWKLFKV	NVCRNHNSY TLSNFNHSKM KEVEEWKKSLGFAPGKG KIQLEEENLE KVEEKLIPED SFQKYPQGKL	KEF.	ADEEEFTIYK ENHKNAIQPT QKILEYLRNK RPTSFTKIYS PTQIADRIKE IVNSDRKTTK LTTNTKLLNS HLMLKTLKNRMFK -DPFGFAVFN DKQKNIK LNLNQILMDS QLVFRNLKDML-G -QKIGYSVFD	K.K LN.NLS QL.LLKN	FKQRLLKKGFN NVLPELYFMKFD VKSCYD YD-DVMKKYE EFVCKWKQVH CPKLFFATMD IEKCYD NK-QISEKFA QFIEKWKNKG RPCLYYVTL
EST2 pep Euplotes pep Trans of tetrahymen	Consensus	EST2 pep Euplotes pep Trans of tetrahymen	Consensus	EST2 pep Euplotes pep Trans of tetrahymen	Consensus	EST2 pep Euplotes pep Trans of tetrahymen



S-1: FFY VTE TTF QKN RLF FYR KSV WSK S-2: RQH LKR VQL RDV SEA EVR QHR EA S-3: ART FRR EKR AER LTS RVK ALF SVL NYE

A-1: AKF LHW LMS VYV VEL LRS FFY VTE TTF Q A-2: LFF YRK SVW SKL QSI GIR QHL KRV QLR DVS A-3: PAL LTS RLR FIP KPD GLR PIV NMD YVV

FIG. 54

D D Y L L I T
3'- ctg ctg atg gag gag tag tgg -5'
a a a a a a a a a
t t t t
c c
Poly 1



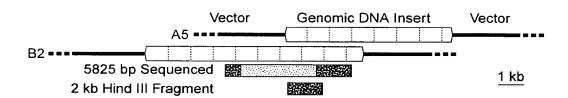


FIG. 55A

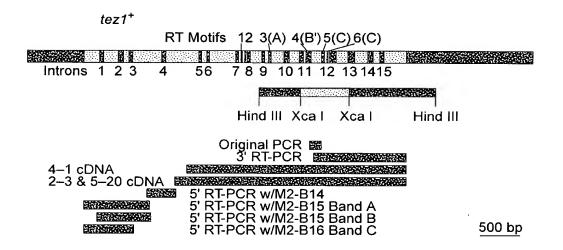
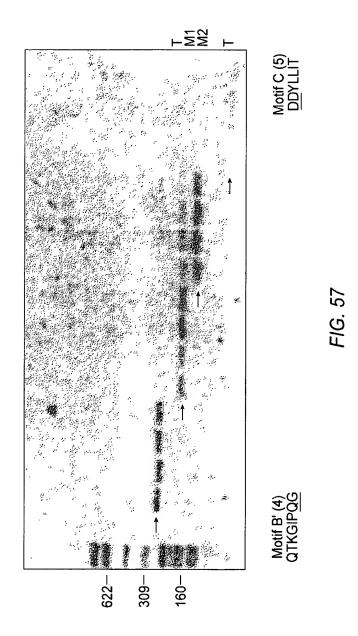


FIG. 55B







DGLFQGSSLSAPIVDLVYDDLLEFYSEFKASPS-----QDTLILKLADDFLIIS KGI PQGLCVSSILSSFYYATLEESSLGFLRDESMNPENPNVNLLMRLTDDYLLIT LCVSYILSSFYYANLEENALQFLRKESMDPEKPETNLLMRLT SILSSFLCHFYMEDLIDEYLSFTKKK------GSVLLRVV Sp_M2 Sc_p103 Ea p123

<---Actual Genomic Sequence. cag gg..... Ö atc caa aaa gtt ggt ტ Ø

Poly 4
t t c
t a a g c c t c g
cag acc aaa gga att cca taa gg

ag acc aaa gga att cca tca ggC TCA ATT CTG TCA TCT TTT TTG TGT CAT TTC TAT ATG tc tgg ttt cct taa ggt agt ccG AGT TAA GAC AGT AGA AAA AAC ACA GTA AAG ATA TAC

FIG. 58A

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GAA GAT TTG ATT GAT GAA TAC CTA TCG TTT ACG AAA AAG AAA GGA TCA GTG TTG TTA CGA CTT CTA AAC TAA CTA ATG GAT AGC AAA TGC TTT TTC TTT CCT AGT CAC AAT GCT

24 ᆸ Ц ß IJ × 团 ы

GTA GTC gac gac tac ctc ctc atc acc CAT CAG ctg ctg atg gag gag tag tgg

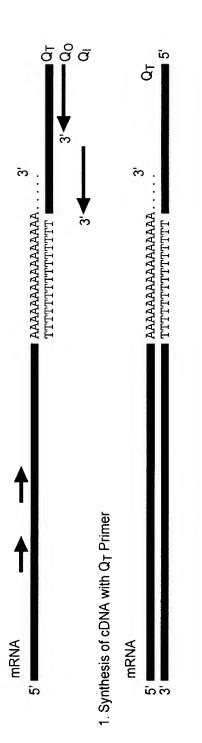
>

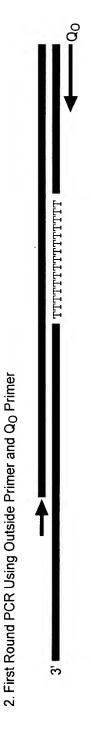
D D Y L L I T
ctg ctg atg gag gag tag tgg
a a a a a a a a a
t t t
c c

<---Actual Genomic Sequencegac gat ttc ctc ttt ata aca...... D D F L F I T

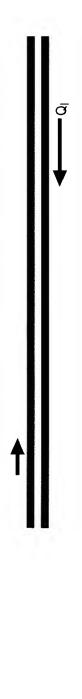
F/G. 58B



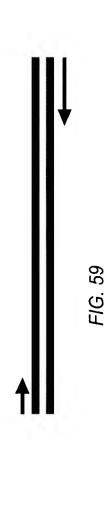




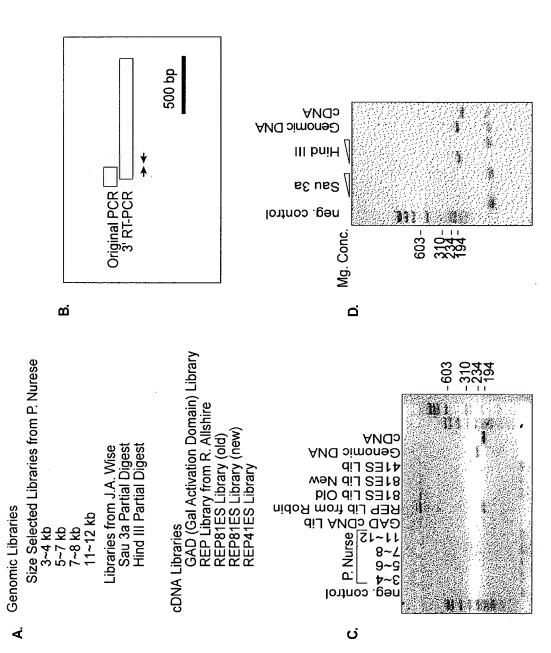
3. Second Round PCR Using Inside Primer and Q_l Primer



4. Sequence Second Round PCR Products Using Inside Primer Q₁ Primer









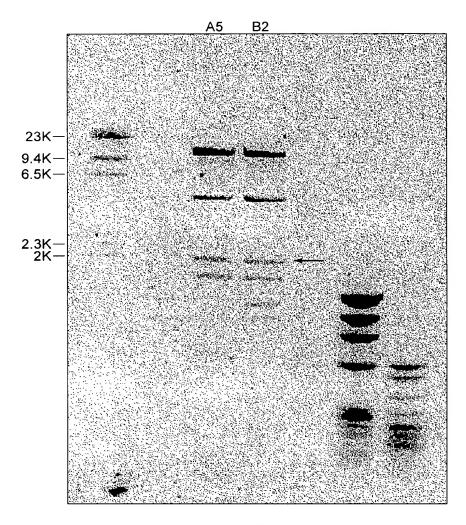
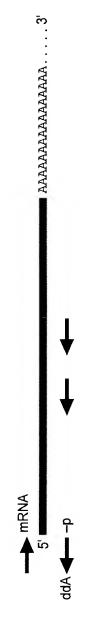
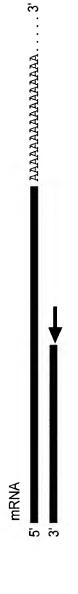


FIG. 61

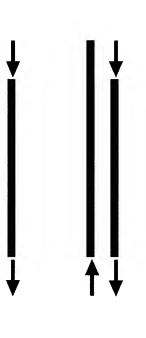




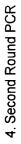
1. Synthesis of cDNA with Specific Destination Primer

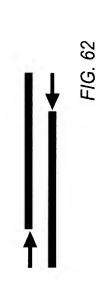


2. Ligate Oligo with 5' -P and blocked 3' to cDNA using T4 RNA Ligase



3. First Round PCR







```
LLMRLTDDYLLITTQENNAVLFIEKLINVSRENGFKFNMKKLQTSFPLS
                                                                                                                                                                                                                                                                                                          VLLRVVDDFLFITVNKKDAKKFLNLSLRGFEKHNFSTSLEKTVINFENS
                                                                                                                                                                                                                                                                                                                      LILKLADDFLIISTDQQQVINIKKLAMGGFQKYNAKANRDKILAVSSQS
                                                                                                                                                                                                                                                                                              cK h
                                                                                                                                                                                                                                                         ... (14) ...
                                                                                                                                                                                                                                                                                  Motif 6(D)
                                                                                                                                                                                                                                ...(9)...
                                                                                                                                                                                                                                             ...(8)...
          WLYNSFIIPILQSFFYITESSDLRNRTVYFRKDIW
                     (366). WLFRQLIPKIIQTFFYCTEISSTVT-IVYFRHDTW
                                 . WIFEDLVVSLIRCFFYVTEQQKSYSKTYYYRKNIW
                                                                                                                                                                                                                                                                                              Gh h
                                                                                                                                                                                 ... (107)...
                                                                                                                                                                       ... (75) ...
                                                                                              ... (62) ...
                                                                                                          ...(61)...
                                                                                                                                                                                                                                YLOKVGIPOGSILSSFLCHFYMEDLIDEYLSF
                                                                                                                                                                                                                                                         YKQTKGIPQGLCVSSILSSFYYATLEESSLGF
                                                                                                                                                                                                                                              YIREDGLFQGSSLSAPIVDLVYDDLLEFYSEF
Motif 0
                                                                                                                                                                                 KLFFATMDIEKCYDSVNREKLSTFLK
                                                                                                                                                          KKYFVRIDIKSCYDRIKQDLMFRIVK
                                                                                                                                                                       ELY FMK FDVKS CYDS I PRMECMRILK
                                                          X
X
                                                                                  AVIRLLPKK--NTFRLITN-LRKRF
                                                                                             SKMRIIPKKSNNEFRIIAIPCRGAD
                                                                                                         GKLRLIPKK--TTFRPIMTFNKKIV
                                                         Motif 2
                                                                                                                                                                                                                    pP hh
                                                                      hR h
                                                                                                                                               Ч
                                                                                                                                                                                                                                                                                 Y Motif 5(C)
                                                                                                                                              ĞΫ
                                                                                                                                  Motif 3(A) AF
                                                                                                                                                                                                        Motif 4(B')
                                                                                                                                                                                                                                                                                              F DDhhh
                                                                                                                                                                                                                      hPQG
                                                                                                                                               hDh
                                                        Motif 1
p hh h K
          (429).
                                   (441)
         Tez1p
Est2p
                                                                                                                                                                                                                                Tez1p
Est2p
                                                                                 Tez1p
Est2p
                                                                                                                                                                                                                                                                                                         Tez1p
Est2p
                                                                                                                                                                      Est2p
                                                                                                                                                           Tez1p
                                 p123
                                                                                                         p123
                                                                                                                                                                                p123
                                                                                                                                                                                                                                                        p123
                                                                                                                                                                                                                                                                                                                                 p123
          SS.Б
Б.С.Б
.
                                                                                                                                                          S S Н
Б С С Б
                                                                                                                                                                                                                                 SS.В
В.С.Б.
```

F/G. 63



155 123 152 88 55 185 218 183 217 122 93 122 90 67 94 24 7 33 57 35 61 _ \ _ _ \ _ _ O O O I ZQI STY SOZ ZIY \bot \bot \vdash Σ • >> • • шци **ω @ Q** 4 . SEL KAV ВSA ⊢ Z ∢ **1** . \times \square \succ PNDNYLQISGIPL - GQFFTQIVGNRC SEGTLVQFCGNNV STFPNYLISILESKNWQLLL CHSANVNVTLLKGAAWKMFH LTALSTQKQYFFQDEWNQVR **О** II II **н**О . $\infty \vdash \times$ N · E **∀** ∟ ♡ > = -OI Z りられ SOF PKCSC . . O IKO **၈** ပ – . 노 Z Y Y RKO ш S Z Z . . 0 • Ш ᄶᄑᆚᅩ . . ~ EGVQFSSP (IALPCLPG IARKEVFS • Ш **ш** > ш **z** – **L** - 4 Z - 4 Z □ 0 □ · 0 <u>U ' '</u> രമര ALKTO SFERRA-NLLMKELYN - NVLTESD VSDRQKLQCF **Z** ' ' œ **م** . ه EHHTPKS $S \vdash \Box$ S A A JZH · 0 : <u>-</u> တ ဟ ဟ ¬×> _ S S _ マ・ス 4 1 . D ٠ ، 🗅 ᅐᅜᇿ スらメ ADYVQLVLRGSP 2DKLDIDLQTN-2KVIRCRNQSQ-SKG SI XXX XXX SSD T T III Z 8 0 0 0 X . X . 0 . Q O – Z > L 」 MHYLLS FVDLL FRHLY > 4 O > A ≻ S T T T <u>ш</u>об **⊢**≷⊼ |>>| ·Z \Box \vdash \circ V V K Q M F C C I V F C L V E L L S EVDVD 0 ' . Q N A N N X **⊢** ∪ **⋈** 日内ロ шаг S D A N T Y A B N E L \bot \bullet >ZOO z > > - マ ПшП ZIJ 34 8 58 36 62 91 68 95 156 124 153 88 88 88 28 28 23 23 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123

F/G. 64/



284 223 275 313 252 308 342 282 341 359 299 374 392 332 406 425 362 437 HHRRLS KKBLN KDFN OF E E X S L C L C L C L C L C - ≥ ш • . 노 XQX JmD ٠ш AN-٠ш SSI O R R - - - V F A F L R S I L V R V F I - - - V L K F I I V I L Q K L L L N K C V T Q F I N E F F Y N I L $\frac{\pi \sigma}{\sigma}$ ZWH IΩZ QFGLINAFQVKQLH SS - - - SIKKLTDLR HEFVSNKNNISAMD - QSTVVPKRLLKVYPLIEQTAKR - LVKIPQRLKVRINLTLQKLLKR RFNRIRKKLKDKVIEKIAYMLEK _ ∽ ∾ ≻ 마 조 조 ᆚᄔᆇ α \neg I F Y R S S Y K K F K Q D L Y E N K Q F L H K L N I N V P N W N N M K S R T R I F E E E SPI X O Z **ㅛ 그 ㅡ __ v _** KLSRYESF SLPLNGYL ELNKHELI SQL **ш** ш **Т** SNAKMCLSD.
DIWFIKHNE
AKHEYYFDH > & z S S III YID-THDDEKILS PLEGTVLDLSHLS PENWRERKQKIE FEILKDLETFLK KGKLIKNLLLS RKNFQKKVKKYVE SI **د** ، ۵ EWLVLGKR RWLFIS - -SWMQVETS SK-FPRONCE ٠ ۵ • · **-**• <u>₽ ₽ ₽</u> ı · **-**S Z · > • 🗕 $\circ \circ \circ$ · 0 J ≻ Z } ' ' 1 14 œ > – S . . OX. - SE T K D F <u>≷</u> S ·I エのメ . _ TNF TNF X-X 1 1 Ω • Ш . . . S I . . • Ш > . . – · > S · **Y** . .Z メタト · > S > N . ш . . のヱ⊢ . 7 Z >⊢ ╙ ZZ **M**·**M** 184 218 252 201 249 285 224 276 314 253 309 343 283 342 360 300 375 333 407 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p p123

FIG. 64E



458 394 470 491 427 503 522 460 534 616 557 630 552 491 564 584 524 597 . В Н Н oσ≥ しゅ> スココ \cdot \vdash \sqsubseteq K L L T F K K D L L K H R M F G R - K I R I K E F K G R L L K K F N N V L P E V M K Y E E F V C K W K Q V G Q P K ΣZW **4** ← - کے Q $\alpha \alpha x$ · 🗠 · шzσ 民田里 . 1 - K X S S S m X K N N M дшσ L Z Q • > **ユコス** <u>スス</u>コ RKD I WKLLCRPF I T SMKMEAFEK I NENRHDTWN KLITPFIVEYFKTYLVENNVC RKNIWDVIMKMSIADLKKETLAEVQEK Ω VSTNQTLRPVASILKHLENHKNAIQPTQKILEYLLTNTKLLNSHLMLKTL ¥ ⊢ ∢ ∑ $\overline{\times}$ \vdash ロるメ Σ <u>s – a</u> IDIKSCYDRIKQDLMFRIVKK FDVKSCYDSIPRMECMRILKD MDIEKCYDSVNREKLSTFLKT • Ш _ <u>_</u> _ _ . 7 шшш 2 2 2 ¥ PILQSFFYITE PKIIQTFFYCTE VSLIRCFFYVTE шшш œ PKKSNNT PKKSNNT PKKSNNT • 14 . . Z Z · 0 G - - - - G R R L I E V Y M Y D D Y Y D D Y YWLYNSFIIP SWLFRQLIPK RWIFEDLVVS Ø⊢Z -51 HZY TTLPPAVI SNFNHSKM SLGEAPGKL $A \vdash \alpha$ Z G > □ ⊢ r IZY **- 4 -**H S A \vdash \vdash \vdash ДХП ——О В У П A > QSI > 0 4 **5** · · スドー メるT \ X \ X X \ X I \ S Y F V F V F M V F > # & ->-8 8 8 SHE $\Sigma \sqcup \Omega$ SHO шш と 目 ら **- ш z** ESK 426 363 438 492 428 504 459 395 471 523 461 535 553 492 565 585 525 598 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123

F/G. 64(



698 624 729 731 657 762 786 713 828 591 696 756 684 795 849 772 894 ОПО ППП ПХП 000 N X - > ₽ | | | **⊘**∪⊩ υZщ • . 4 . 4 . I VK - GNSQ TALWVEDKO IY - NFNGKF . 7 • VNMRSLDT ELEVWKHSS DMKTLAL _ F X C · 0 **ス**Q E ZOL F & し マンド ストス ストス SSS N T T T ZØJ FLCHFYMEDLID PIVDLVYDDLLE ILSSFYYATLEE шσд TKHMGKSFF ALFNTRIS 2TKKASMWL LNLSLRGFEKHNFSTSLEKTVINF KKLAMGGFQKYNAKANRDKILAVS IEKLINVSRENGFKFNMKKLQTSF MELLSGHI MELFKTA NICQYN) . .Z п Х — О П О - S V L L R V V D D F L F - T L I L K L A D D F L I V N L L M R L T D D Y L L DMVPFEKVVQLLS
-VLKLFNVVNASR
QYPTLFSVLENEQI KKRMPFFGF --MHIEVK DYCDWIGI X N Q M > q N > y X M -S N N N > K J S S S _ > _ _ _ _ _ \times \cdot \circ S L Z GMDSVEEQNIVA ST 3 2 2 2 2 2 1 ZYO SSL . .Z н S I · (D <u>o</u> o <u>o</u> ٠ ۵ _ v _ · (5) > aaa ·Z マド ら > - X × - X × - X × - X ഗ - Ш ΔЩΔ • ш ШОШ . _ ш . . . EA · OZ · 02 $\overline{\Delta \pm}$ \neg S . 7 SOS . Q $\Delta \perp Z$ > · 0 · ·> ᄌᄑᇳ OZш · L よらり **II** - II · **Y VZZ** 935 571 564 566 592 597 699 625 730 732 658 763 757 685 796 714 829 817 740 862 787 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123

FIG. 64[



915 821 960 793 927 AV-GRK-WKK ASFLQR--EM EVSK--EM AY IY SDQC · × ~ SZL MCM a o i · > . GG X O X - N G O III O LAEILGYTSRRFLSSAEVKWLFCLGMRRTVSGCPITKCDPLIEYEVRFTILNGFLTRAFFKYLVCNIKDTIFGEEHYPDFFL L L N O F H S AQAYLKRMKDIFIPQRMFITD ---YKSAFKDLSIN --VTQNM YMQCAKEYKDHFKKNLAMSSM A > > $\neg \sigma$ · S Ω. 8 – 7 8 ± – I • ш . — > _ H ≻ Q KS田 ____ \bot \vdash -855 994 850 773 895 883 794 928 916 822 961 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123

-1G. 64E



122 93 122 155 123 152 188 155 185 218 183 217 57 35 61 90 67 94 24 7 33 **ШЧХ** >>> Ω⊢Х ⊢∟≥ SOZ ZIY I 000 ZGI O E O **400** \bot \bot \vdash Σ . . > -ドョロ **ω © Q 4** ' ' **- 」** 2 μΖц ⊢ Z ∢ ביי M S A しい> \times \mathbb{H} \succ g н н ШАШ **コエ** R O K Z FD . $\alpha \vdash x$ > = υшα **」** └ > – z z のメー A L O Q M M W Z Z 0 0 0 りらく **GIZ** SOL တ ပ — S S SHRG ' \ Ш RKO **℃** ∟ ७ $Z \not\subseteq Y$ **—** — 止 1 10 _ _ ш メロト **∑** – ∟ ZKU ggg 1 10 **F** ➤ 〒 S ᄎ Ω トェイトン ш • Ш $\alpha z \rightarrow$ പ വ വ . . . ιШ ШШО യമം z – L NYLISILES NNVNVTLLKG STQKQYFFQ S _ > **000 スルト** . 0 **U** ' ' ggш RR - N L L M KG - . · |-шОШ \Box \Box \Box $\vdash \forall z$ P N I · **Y Z** ' ' **- ∢ ∑** œ のメメ S . _ $S \vdash D$ <u>ዋ</u> አ . 4 \succ JZH - < < · S ш — — SHI BSA ロド> ·I လ လ လ — თ ტ SKGS-FE-NYTV-G-TKYL-FG I マ・ス **⊢** ഗ ഗ KRK VN · N Ш . Q **د** ب 🗅 ᅐᇝᅚ L A D R S Y ·I $\sigma z \sigma$ တတ LSA Y S G T T III Σ **ე** ⊢ თ ·Z $\vdash \bot \vdash$ $\kappa \kappa \star$ N Q Q . . 0 SOL ¥ > V × × SE . 4 メ・ス > エ 」 ACRICATION OF THE PROPERTY OF HY COLLS · 0 **コーエ** SQD ШGS ·Z > 4 0 **□** ∽ $\Box \vdash \varnothing$. 0 Z - - -**下** C R ZZY · > > _ - \circ \vdash \circ OZ < 国内口 ≻⊻> шФZ . 0 $I \vdash \vdash$ gγш >> _ Σ ਘ ਘ C C K Т — > А L L · > $\circ \circ \times$ **O** · · A A J **╙** � > Z : : S L Z zgg · W > 」と — ш — ZIJ 25 8 34 58 36 62 91 68 95 156 124 153 86 156 88 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123

FIG. 641



284 223 275 251 200 248 313 252 308 342 282 341 359 299 374 392 332 406 425 362 437 _ = □ - = □ _ & _ σzz • ' ' **4** – ≥ **4 4 ⊅** ОЕЕ **- - 4** ٠Ш u ∢ - - > ールエ **–** – σ 2 2 2 スロス • • Ш > 4 4 444 SSI エスス aa> Q R R ᆂᄪᇠ **コエ>** ш___ Z W H T S C IΩZ $\times z -$ DLH DLR AMDR アスス しのト > _ _ $\kappa \kappa \star$ **し F 又** K X Z トイ 日 SOI $ZZ \bot$ > o > ᄔᆜᅳ шшш \times \vdash \circ QX≻ ⊢¬∑ > Z & SQL __ _ _ __ S > _ шшт ___ > _ _ > ZZ **—** — **ш** ШΦШ NAKMCLSD DIWFTKHN AKHFYYFD A – X F X S Q X S D - THDDEK I LS. EGTVLDLSHLSI ENWRERKQK I EI ШQК X L X りドド **S** > Ш SSYKKFKQD ---KQFLHK NWNNMKSR 기 — — R — N S L Z ZOZ H I K D L E T F L K L K I I K N L N L L S L N F Q K V K K Y V E L _ · \overline{\sigma} **┙・>** ு ட S . . μош ٠ ۵ . . . S OOI V L G K D - S - C D - E - S \propto · > スSY • ·Z <u>≻</u> ·Z σ σ imes. 0 • SK-FF NOFF 7 T T _ P _ _ . — • ·Z Z Z Z S ·Z Φ Φ Φ · 0 $\circ \circ \circ$ m <u>♀</u> <u>≺</u> . 1 1 11 œ > - s 国民の . _ ・エ エペス S ᄦᅑᅈ — ш · 0 ZZH • Ш **-zz** ШОШ \ \ \ \ \ ഗ $x \times x$ I • Ш らドR > . . . — QK . - **-** -· > . 7 \vdash . .Z S . . メット · > z v v $\times \times z$ ¥G FG . . Z oz⊢ $\sigma > z$ · × _ _ _ **α** · **α** $\perp > \perp$ >⊢ ╙ $z \times x$ 249 285 224 276 314 253 309 283 184 252 201 343 360 300 393 333 407 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123

⁻1G. 64(



458 394 470 491 427 503 522 460 534 552 491 564 616 557 630 634 570 663 584 524 597 ۵ v ≷ **→** □ > шш \cdot \vdash \vdash **メ** 」 」 ・エ ᄌᇚᄌ ZZW L V -. ₾ ≥ . O D Ø 202 . 44 民田田 . 22 . ШZσ >z> **ス K X** 日内民 **R** - 0 дшо Z — F R O Z FKKDLLKHRMFG FKQRLLKKFNNV YEEFVCKWKQVG ZKW ZZZ ZUY ススー . 0 $\neg \vdash \circ$ ш > ш **レース** . 7 ⊢∢⋝ エント ロのメ ZZO メタト . ≥ 8 S S -z> _ _ _ メロト • Ш $\omega - \sigma$ \mathbf{z} \mathbf{m} \mathbf{m} ススス ' \ шшш **Ⅲ > ∢** 2 2 2 · × шшш **4** - 1 - 1 \vdash らドー **—** — **ц** · 🗠 NT LRPVAS IA I QPT QK KLLNSHL A ≻ F - c> - Ш - $\alpha \alpha \vdash$ L ≥ ω ZZH Z ⊼ π ⊼ ∑ ⊼ ⊼ ' Z ' • 🔀 ·S · W Z S ⊢ ∪ **ススス** ススス と・0 $\vdash \square \prec$ ィースス O R R P L C F > A ス R > ㄱ - 둘 - - > 주 <u>S</u> 도 **____** ØZ⊢ ____ ZYZ E V Y M Y N Y D D V \vdash \bot \vdash α SF-F QC-PF DCVVS SZH ZH ح ۵ ≥ $\sigma \vdash z$ KLCR VLCR >>> 000 $\vdash Z Y$ コスス $A \vdash \alpha$ L P P A V ≥≻⊢ らられ IZY \neg \vdash \vdash YWLYNS SWLFRO RWLFRO $\mathbf{x} - \mathbf{\vdash}$ z 4 > ススヨ **- ㅛ 그** - F A >>> >>> H O A $\vdash \bot \vdash$ — — Q Ф ≻ щ <>O トz (5 SI > O 4 **⊢** ⊗ **⊣ O** · · ΩХπ $\mathbf{K} \times \mathbf{H}$ $\times \circ \vdash$ ≥ w ∩ SHE スエス としと > ≥ < $\alpha \alpha \Sigma$ ⊢≻⊼ Ω⊢⊼ ᄪᄪᅩ \mathbf{x} \mathbf{x} \rightarrow \rightarrow \perp ド 三 ら SED ->-至らよ $- \mathbf{W} \mathbf{z}$ 国 ら K > ╙ ≩ 363 438 459 395 471 492 428 504 523 461 535 553 492 565 585 525 598 617 558 631 Tip1p Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123

FIG. 64F



591 696 698 624 729 731 657 762 756 684 795 786 713 828 739 861 849 772 894 | | | | | | | go⊩ ≻≻≻ \times \times J∑ G YZJ ı . 4 SZL トメー · 1 $\exists \vdash \Xi$ SHS のメメ шшш **⊢** ഗ ¬ . . . R ロ X - Ш -ZOU <u>о ш о</u> **5** · 0 DOA MRSLE VWKHS S d X ロ > X OШZ ר ט ר YØZ ZOL **-- -- ×** 즈 호교 — ¥ S S × ≺ ススス လ လ လ $\overline{\Sigma}$ ZØJ ZOG \Box \Box \Box > - шσд FSL $\vdash \circ \vdash$ μоц ZWO **L** – ≥ ۵ ۰ IHZ $z > \omega$ > _ _ SAS KHMGKS ALFNTF VTKKASN T V | | . .Z ロドト တ မာ တ CHFYMED: LCHFYMED: LSSFYYAT MPFFGF MH FVKE CDW G - 8 1 - S SEG EKVQLLS FNVVNASR FSVLENEG STSLEK KANRDK KFNMKK エШー _ _ Z Z _ _ Z **∝** · ≻ ШSZ X . Q ш Ф — > と し Р Х Р Г Л Л -> -- - -SAS **の** L Z **ω** ∢ > တ တ တ N N N ШOZ S 5 ⊢ SOD _ თ ე ZYU SSX ш G № m a a · 0 ഗഗച ٠Z ト S L 000 · (5 ٠ م FFS _ ი _ · (5 ス し F $\alpha\alpha\alpha$ ტ ტ > ·Z タメら • Ш _ L _ S шοш • Ш M O Z Щ . . _ **⊿ ∑ −** ZOS 0 K -- 0 0 - • **∑** ⋖ · 4 · 0 Z SAI **--** ≃ ш . — DKQ <u>> 0 X</u> · 02 ト し と スエー . . 0 >>× ス lu lu S ・マ **5** 0 0 ᆂ $\Delta \perp Z$ LZ4 o k o > · 0 **ーメー** メ P 田 > OZ-メるロ **ட** – ட **VZZ** 699 625 730 732 658 763 571 664 666 592 697 757 585 796 787 714 829 817 740 862 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123

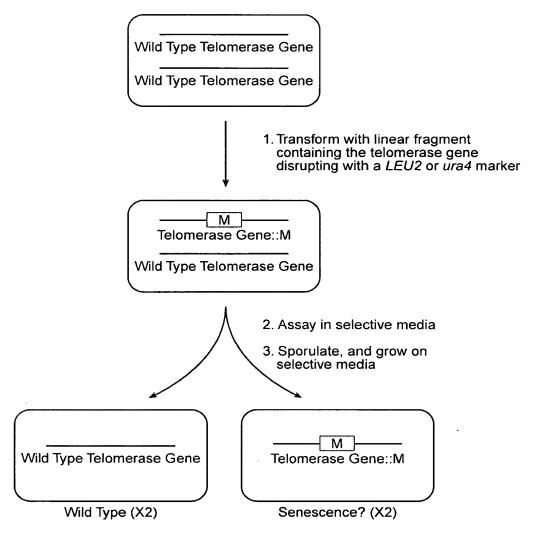
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915 821 960 981 877 988 884 1031 882 793 927 > - - -> - - -> = - -RQVLF QAYLF SDQC • 🔀 \propto SZL ≥ L O I • > SMC · (5) **ス**のス · (J L L · 0 <u>ი</u> ა ⊢ OLIKPLRPVLR LLRKEIQHLC 11LKAKEAKLK IVIGRE ISFLOR EVSKI O M S G $\mathbf{Z} - \mathbf{J}$ ე ე ⊩ ≅ ⊩ ⊩ , **X** SCCN | YRL | SEC - - - -TOLLN TONMOFHS ASSMIDLE (WLFCL(RFTILN(EHYPDF - - KFKDN--L KKY-FNRVCM-ᇫᅂᇤ ORMF--VATOR **z** – 4 >> ७ LZO шшш STAQVF-D-THNSK STNTVLMQ-DHVVK-NN-THYFRKT-TTE <≻ωш⊢ QAYLKRMKD - F - P - - - YKSAFKDLS - N MQCAKEYKDHFKK თ **–** വ **ユース** ш Ф — LGYTSRRF SCPITKCDP FKYLVCNI , X $\sigma \cdot \vdash$ ш . . ш . . **O** 1 1 <>≻ **D** · S — ტ **ட** I . . $\alpha = -$ <u>~ - -</u> шoд > . — H - S **ζZ**Σ ∢>℃ KSE **∢**·≻ 916 822 961 949 855 994 883 794 928 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 Sp_Tip1p Sc_Est2p Ea_p123 മ്

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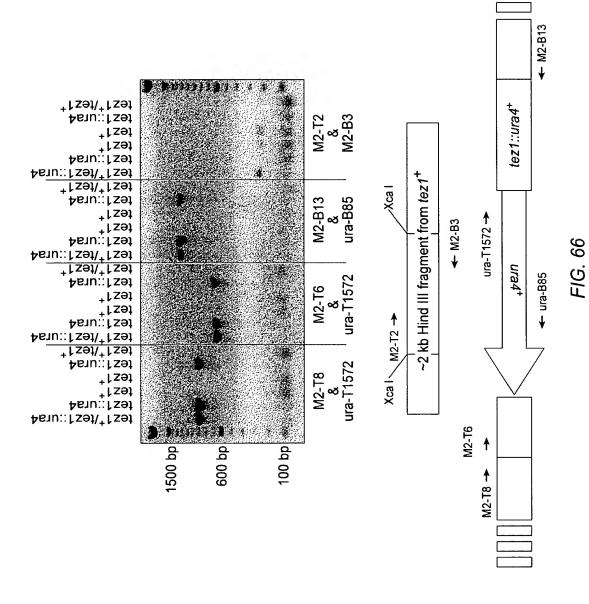




(These cells will show a senescence phenotype if the disrupted gene encodes a telomerase subunit.)

FIG. 65





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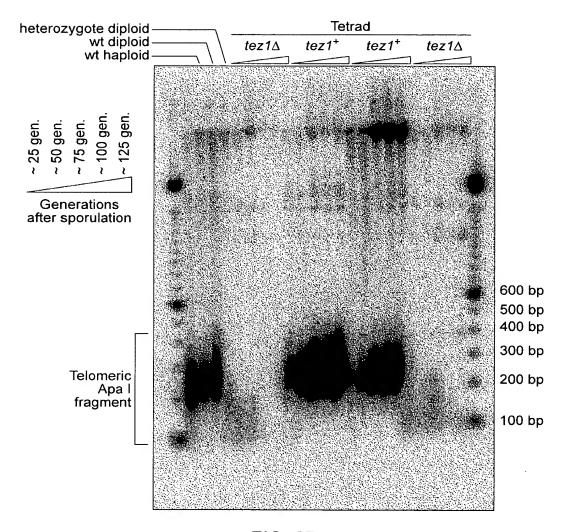


FIG. 67



met ser val tyr val val glu leu leu GCCAAGTTCCTGCACTGGCTG ATG AGT GTG TAC GTC GAG CTG CTC 1.0 arg ser phe phe tyr val thr glu thr thr phe gln lys asn arg AGG TCT TTC TTT TAT GTC ACG GAG ACC ACG TTT CAA AAG AAC AGG 30 leu phe phe tyr arg lys ser val trp ser lys leu gln ser ile CTC TTT TTC TAC CGG AAG AGT GTC TGG AGC AAG TTG CAA AGC ATT gly ile arg gln his leu lys arg val gln leu arg glu leu ser GGA ATC AGA CAG CAC TTG AAG AGG GTG CAG CTG CGG GAG CTG TCG glu ala glu val arg gln his arg glu ala arg pro ala leu leu GAA GCA GAG GTC AGG CAG CAT CGG GAA GCC AGG CCC GCC CTG CTG 80 thr ser arg leu arg phe ile pro lys pro asp gly leu arg pro ACG TCC AGA CTC CGC TTC ATC CCC AAG CCT GAC GGG CTG CGG CCG 90 ile val asn met asp tyr val val gly ala arg thr phe arg arg ATT GTG AAC ATG GAC TAC GTC GTG GGA GCC AGA ACG TTC CGC AGA 100 110 glu lys ala glu arg leu thr ser arg val lys ala leu phe GAA AAG ARG GCC GAG CGT CTC ACC TCG AGG GTG AAG GCA CTG TTC 120 ser val leu asn tyr glu arg ala arg arg pro gly leu leu gly AGC GTG CTC AAC TAC GAG CGG GCG CGC CGC CCC GGC CTC CTG GGC 140 ala ser val leu gly leu asp asp ile his arg ala trp arg thr GCC TCT GTG CTG GGC CTG GAC GAT ATC CAC AGG GCC TGG CGC ACC 150 phe val leu arg val arg ala gln asp pro pro pro glu leu tyr TTC GTG CTG CGT GTG CGG GCC CAG GAC CCG CCG CCT GAG CTG TAC phe val lys val asp val thr gly ala tyr asp thr ile pro gln TTT GTC AAG GTG GAT GTG ACG GGC GCG TAC GAC ACC ATC CCC CAG asp arg leu thr glu val ile ala ser ile ile lys pro gln asn GAC AGG CTC ACG GAG GTC ATC GCC AGC ATC ATC AAA CCC CAG AAC thr tyr cys val arg arg tyr ala val val gln lys ala ala met ACG TAC TGC GTG CGT CGG TAT GCC GTG GTC CAG AAG GCC GCC ATG

FIG. 68A



210 gly thr ser ala arg pro ser arg ala thr ser tyr val gln cys ĞGC ACG TCC GCA AGĞ CCT TCA AGĂ GCC ACG TCC TÂC GTC CAG TGC 230 220 gln gly ile pro gln gly ser ile leu ser thr leu leu cys ser CAG GGG ATC CCG CAG GGC TCC ATC CTC TCC ACG CTG CTC TGC AGC 240 leu cys tyr gly asp met glu asn lys leu phe ala gly ile arg CTG TGC TAC GGC GAC ATG GAG AAC AAG CTG TTT GCG GGG ATT CGG 260 arg asp gly leu leu leu arg leu val asp asp phe leu leu val CGG GAC GGG CTG CTC CTG CGT TTG GTG GAT GAT TTC TTG TTG GTG thr pro his leu thr his ala lys thr phe leu arg thr leu val ACA CCT CAC CTC ACC CAC GCG AAA ACC TTC CTC AGG ACC CTG GTC 290 arg gly val pro glu tyr gly cys val val asn leu arg lys thr CGA GGT GTC CCT GAG TAT GGC TGC GTG GTG AAC TTG CGG AAG ACA 300 val val asn phe pro val glu asp glu ala leu gly gly thr ala GTG GTG AAC TTC CCT GTA GAA GAC GAG GCC CTG GGT GGC ACG GCT 310 phe val gln met pro ala his gly leu phe pro trp cys gly leu TTT GTT CAG ATG CCG GCC CAC GGC CTA TTC CCC TGG TGC GGC CTG 330 leu leu asp thr arg thr leu glu val gln ser asp tyr ser ser CTG CTG GAT ACC CGG ACC CTG GAG GTG CAG AGC GAC TAC TCC AGC tyr ala arg thr ser ile arg ala ser leu thr phe asn arg gly TÂT GCC CGG ACC TCC ATC AGA GCC AGT CTC ACC TTC AAC CGC GGC 360 phe lys ala gly arg asn met arg arg lys leu phe gly val leu TTC AAG GCT GGG AGG AAC ATG CGT CGC AAA CTC TTT GGG GTC TTG arg leu lys cys his ser leu phe leu asp leu gln val asn ser CGG CTG AAG TGT CAC AGC CTG TTT CTG GAT TTG CAG GTG AAC AGC 390 leu gln thr val cys thr asn ile tyr lys ile leu leu leu gln CTC CAG ACG GTG TGC ACC AAC ATC TAC AAG ATC CTC CTG CAG ala tyr arg phe his ala cys val leu gln leu pro phe his gln GCG TAC AGG TTT CAC GCA TGT GTG CTG CAG CTC CCA TTT CAT CAG

FIG. 68B



420 gln val trp lys asn pro his phe ser cys ala ser ser leu thr CAA GTT TGG AAG AAC CCA CAT TTT TCC TGC GCG TCA TCT CTG ACA 430 440 arg leu pro leu leu leu his pro glu ser gln glu arg arg asp CGG CTC CCT CTG CTA CTC CAT CCT GAA AGC CAA GAA CGC AGG GAT 450 val ala gly gly gln gly arg arg pro ser ala leu arg gly GTC GCT GGG GGC CAA GGG CGC CGC CGG CCC TCT GCC CTC CGA GGC 470 arg ala val ala val pro pro ser ile pro ala gln ala asp ser CGT GCA GTG GCT GTG CCA CCA AGC ATT CCT GCT CAA GCT GAC TCG 480 thr pro cys his leu arg ala thr pro gly val thr gln asp ser ACA CCG TGT CAC CTA CGT GCC ACT CCT GGG GTC ACT CAG GAC AGC 500 pro asp ala ala glu ser glu ala pro gly asp asp ala asp cys CCA GAC GCA GCT GAG TCG GAA GCT CCC GGG GAC GAC GCT GAC TGC pro gly gly arg ser gln pro gly thr ala leu arg leu gln asp CCT GGA GGC CGC AGC CAA CCC GGC ACT GCC CTC AGA CTT CAA GAC his pro gly leu met ala thr arg pro gln pro gly arg glu gln CAT CCT GGA CTG ATG GCC ACC CGC CCA CAG CCA GGC CGA GAG CAG thr pro ala ala leu ser arg arg ala tyr thr ser gln gly gly ACA CCA GCA GCC CTG TCA CGC CGG GCT TAT ACG TCC CAG GGA GGG 550 arg gly gly pro his pro gly leu his arg trp glu ser glu ala AGG GGC GGC CCA CAC CCA GGC CTG CAC CGC TGG GAG TCT GAG GCC 564 ΟP TGA GTGAGTGTTTGGCCGAGGCCTGCATGTCCGGCTGAAGGCTGAGTGTCCGGCTGAGGC ${\sf CTGAGCGAGTGTCCAGCCAAGGGCTGAGTGTCCAGCACCTGCGTTTTCACTTCCCCAC}$ AGGCTGGCGTTCGGTCCACCCCAGGGCCAGCTTTTCCTCACCAGGAGCCCGGCTTCCACT CCCCACATAGGAATAGTCCATCCCCAGATTCGCCATTGTTCACCCTTCGCCCTGCCTTCC TTTGCCTTCCACCCCCACCATTCAGGTGGAGACCCTGAGAAGGACCCTGGGAGCTTTGGG AATTTGGAGTGACCAAAGGTGTGCCCTGTACACAGGCGAGGACCCTGCACCTGGATGGGG GTCCCTGTGGGTCAAATTGGGGGGGGGTGCTGTGGGAGTAAAATACTGAATATATGAGTT TTTCAGTTTTGGAAAAAAAAAAAAAAAAAAAAAAAAA

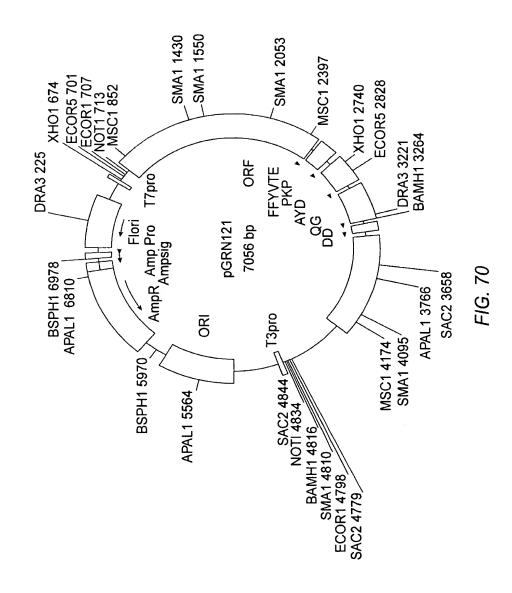
FIG. 68C



```
Motif -1
               ...LVVSLIRCFFYVTEQQKSYSKT...
Ep p123
Sp Tez1
               ...FIIPILQSFFYITESSDLRNRT...
Sc Est2
               ...LIPKIIQTFFYCTEISSTVTIV...
Hs TCP1
               ...YVVELLRSFFYVTETTFQKNRL...
consensus
                           FFY TE
                                                   K
Motif 0
                          p hhh K
                                       hR h
                                                    R
Ep p123
Sp Tez1
               ... KSLGFAPGKLRLIPKKT--TFRPIMTFNKKIV...
               ...QKTTLPPAVIRLLPKKN--TFRLITNLRKRFL...
Sc Est2
               ...TLSNFNHSKMRIIPKKSNNEFRIIAIPCRGAD...
Hs TCP1
               ... ARPALLTSRLRFIPKPD--GLRPIVNMDYVVG...
consensus
                             R PK
                                         R I
                               AF
Motif A
                      h
                         hDh
                              GY
                                   h
Ep p123
Sp Tez1
               ... PKLFFATMDIEKCYDSVNREKLSTFLK...
               ... RKKYFVRIDIKSCYDRIKQDLMFRIVK...
Sc Est2
               ...PELYFMKFDVKSCYDSIPRMECMRILK...
Hs TCP1
               ...PELYFVKVDVTGAYDTIPQDRLTEVIA...//...
consensus
                      F
                          D
Motif B
                             hPQG
                                    pS hh
               ...NGKFYKQTKGIPQGLCVSSILSSFYYA...
Ep p123
Sp Tez1
Sc Est2
               ... GNSQYLQKVGIPQGSILSSFLCHFYME...
               ... EDKCYIREDGLFQGSSLSAPIVDLVYD...
Hs TCP1
               ... RATSYVQCQGIPQGSILSTLLCSLCYG...
consensus
                            G QG
                                    S
                           Υ
Motif C
                       h F DD hhh
Ep p123
               ... PNVNLLMRLTDDYLLITTQENN...
Sp Tez1
Sc Est2
               ...KKGSVLLRVVDDFLFITVNKKD...
               ...SQDTLILKLADDFLIISTDQQQ...
Hs TCP1
               ... RRDGLLLRLVDDFLLVTPHLTH...
consensus
                             DD L
Motif D
                         Gh h cK
               ...NVSRENGFKFNMKKL...
Ep p123
Sp Tez1
               ...LNLSLRGFEKHNFST...
               ...KKLAMGGFQKYNAKA...
Sc Est2
Hs TCP1
               ...LRTLVRGVPEYGCVV...
consensus
                        G
```

FIG. 69







GCAGCGCTGC GTCCTGCTGC GCACGTGGGA AGCCCTGGCC CCGGCCACCC 1 CCGCGATGCC GCGCGCTCCC CGCTGCCGAG CCGTGCGCTC CCTGCTGCGC 51 AGCCACTACC GCGAGGTGCT GCCGCTGGCC ACGTTCGTGC GGCGCCTGGG 101 GCCCAGGGC TGGCGGCTGG TGCAGCGCGG GGACCCGGCG GCTTTCCGCG 151 CGNTGGTGGC CCANTGCNTG GTGTGCGTGC CCTGGGANGN ANGGCNGCCC 201 CCCGCCGCCC CCTCCTTCCG CCAGGTGTCC TGCCTGAANG ANCTGGTGGC 251 CCGAGTGCTG CANANGCTGT GCGANCGCGG CGCGAANAAC GTGCTGGCCT 301 351 TCGGCTTCGC GCTGCTGGAC GGGGCCCGCG GGGGCCCCCC CGAGGCCTTC ACCACCAGCG TGCGCAGCTA CCTGCCCAAC ACGGTGACCG ACGCACTGCG 401 GGGGAGCGGG GCGTGGGGC TGCTGCTGCG CCGCGTGGGC GACGACGTGC 451 TGGTTCACCT GCTGGCACGC TGCGCGNTNT TTGTGCTGGT GGNTCCCAGC 501 TGCGCCTACC ANGTGTGCGG GCCGCCGCTG TACCAGCTCG GCGCTGCNAC 551 TCAGGCCGG CCCCGCCAC ACGCTANTGG ACCCGAANGC GTCTGGGATC 601 CAACGGCCT GGAACCATAG CGTCAGGGAG GCCGGGGTCC CCCTGGGCTG 651 CCAGCCCGG GTGCGAGGAG GCGCGGGGGC AGTGCCAGCC GAAGTCTGCC 701 GTTGCCCAAG AGGCCCAGGC GTGGCGCTGC CCCTGAGCCG GAGCGGACGC 751 CCGTTGGGCA GGGGTCCTGG GCCCACCCGG GCAGGACGCC TGGACCGAGT 801 GACCGTGGTT TCTGTGTGGT GTCACCTGCC AGACCCGCCG AAGAAGCCAC 851 CTCTTTGGAG GGTGCGCTCT CTGGCACGCG CCACTCCCAC CCATCCGTGG 901 GCCGCCAGCA CCACGCGGC CCCCCATCCA CATCGCGGCC ACCACGTCCT 951 1001 GGGACACGCC TTGTCCCCCG GTGTACGCCG AGACCAAGCA CTTCCTCTAC TCCTCAGGCG ACAAGNACAC TGCGNCCCTC CTTCCTACTC AATATATCTG 1051 AGGCCCAGCC TGACTGGCGT TCGGGAGGTT CGTGGAGACA NTCTTTCTGG 1101 TTCCAGGCCT TGGATGCCAG GATTCCCCGC AGGTTGCCCC GCCTGCCCCA 1151 1201 GCGNTACTGG CAAATGCGGC CCCTGTTTCT GGAGCTGCTT GGGAACCACG CGCAGTGCCC CTACGGGGTG TTCCTCAAGA CGCACTGCCC GCTGCGAGCT 1251 GCGGTCACCC CAGCAGCCGG TGTCTGTGCC CGGGAGAAGC CCCAGGGCTC 1301 1351 TGTGGCGGCC CCCGAGGAGG AGGAACACAG ACCCCCGTCG CCTGGTGCAG 1401 CTGCTCCGCC AGCACAGCAG CCCCTGGCAG GTGTACGGCT TCGTGCGGGC CTGCCTGCGC CGGCTGGTGC CCCCAGGCCT CTGGGGCTCC AGGCACAACG 1451 AACGCCGCTT CCTCAGGAAC ACCAAGAAGT TCATCTCCCT GGGGAAGCAT 1501 GCCAAGCTCT CGCTGCAGGA GCTGACGTGG AAGATGAGCG TGCGGGACTG 1551 CGCTTGGCTG CGCAGGAGCC CAGGGGTTGG CTGTGTTCCG GCCGCAGAGC 1601 1651 ACCGTCTGCG TGAGGAGATC CTGGCCAAGT TCCTGCACTG GCTGATGAGT GTGTACGTCG TCGAGCTGCT CAGGTCTTTC TTTTATGTCA CGGAGACCAC 1701 GTTTCAAAAG AACAGGCTCT TTTTCTACCG GAAGAGTGTC TGGAGCAAGT 1751 1801 TGCAAAGCAT TGGAATCAGA CAGCACTTGA AGAGGGTGCA GCTGCGGGAG 1851 CTGTCGGAAG CAGAGGTCAG GCAGCATCGG GAAGCCAGGC CCGCCCTGCT GACGTCCAGA CTCCGCTTCA TCCCCAAGCC TGACGGGCTG CGGCCGATTG 1901 1951 TGAACATGGA CTACGTCGTG GGAGCCAGAA CGTTCCGCAG AGAAAAGAGG GCCGAGCGTC TCACCTCGAG GGTGAAGGCA CTGTTCAGCG TGCTCAACTA 2001 2051 CGAGCGGCC CGCCCCCG GCCTCTGGG CGCCTCTGTG CTGGGCCTGG ACGATATCCA CAGGGCCTGG CGCACCTTCG TGCTGCGTGT GCGGGCCCAG 2101 GACCCGCCGC CTGAGCTGTA CTTTGTCAAG GTGGATGTGA CGGGCGCGTA 2151 2201 CGACACCATC CCCCAGGACA GGCTCACGGA GGTCATCGCC AGCATCATCA AACCCAGAA CACGTACTGC GTGCGTCGGT ATGCCGTGGT CCAGAAGGCC 2251 GCCCATGGGC ACGTCCGCAA GGCCTTCAAG AGCCACGTCT CTACCTTGAC 2301 AGACCTCCAG CCGTACATGC GACAGTTCGT GGCTCACCTG CAGGANAACA 2351 GCCCGCTGAG GGATGCCGTC GTCATCGAGC AGAGCTCCTC CCTGAATGAG 2401 2451 GCCAGCAGTG GCCTCTTCGA CGTCTTCCTA CGCTTCATGT GCCACCACGC

FIG. 71A



2501	CGTGCGCATC	AGGGGCAAGT	CCTACGTCCA	GTGCCAGGGG	ATCCCGCAGG
2551	GCTCCATCCT	CTCCACGCTG	CTCTGCAGCC	TGTGCTACGG	CGACATGGAG
2601	AACAAGCTGT	TTGCGGGGAT	TCGGCGGGAC	GGGCTGCTCC	TGCGTTTGGT
2651	GGATGATTTC	TTGTTGGTGA	CACCTCACCT	CACCCACGCG	AAAACCTTCC
2701	TCAGGACCCT	GGTCCGAGGT	GTCCCTGAGT	ATGGCTGCGT	GGTGAACTTG
2751	CGGAAGACAG	TGGTGAACTT	CCCTGTAGAA	GACGAGGCCC	TGGGTGGCAC
2801	GGCTTTTGTT	CAGATGCCGG	CCCACGGCCT	ATTCCCCTGG	TGCGGCCTGC
2851	TGCTGGATAC	CCGGACCCTG	GAGGTGCAGA	GCGACTACTC	CAGCTATGCC
2901	CGGACCTCCA	TCAGAGCCAG	TCTCACCTTC	AACCGCGGCT	TCAAGGCTGG
2951	GAGGAACATG	CGTCGCAAAC	TCTTTGGGGT	CTTGCGGCTG	AAGTGTCACA
3001	GCCTGTTTCT	GGATTTGCAG	GTGAACAGCC	TCCAGACGGT	GTGCACCAAC
3051	ATCTACAAGA	TCCTCCTGCT	GCAGGCGTAC	AGGTTTCACG	CATGTGTGCT
3101	GCAGCTCCCA	TTTCATCAGC	AAGTTTGGAA	GAACCCCACA	TTTTTCCTGC
3151	GCGTCATCTC	TGACACGGCC	TCCCTCTGCT	ACTCCATCCT	GAAAGCCAAG
3201	AACGCAGGGA	TGTCGCTGGG	GGCCAAGGGC	GCCGCCGGCC	CTCTGCCCTC
3251	CGAGGCCGTG	CAGTGGCTGT	GCCACCAAGC	ATTCCTGCTC	AAGCTGACTC
3301	GACACCGTGT	CACCTACGTG	CCACTCCTGG	GGTCACTCAG	GACAGCCCAG
3351	ACGCAGCTGA	GTCGGAAGCT	CCCGGGGACG	ACGCTGACTG	CCCTGGAGGC
3401	CGCAGCCAAC	CCGGCACTGC	CCTCAGACTT	CAAGACCATC	CTGGACTGAT
3451	GGCCACCCGC	CCACAGCCAG	GCCGAGAGCA	GACACCAGCA	GCCCTGTCAC
3501	GCCGGGCTCT	ACGTCCCAGG	GAGGGAGGGG	CGGCCCACAC	CCAGGCCCGC
3551	ACCGCTGGGA	GTCTGAGGCC	TGAGTGAGTG	TTTGGCCGAG	GCCTGCATGT
3601	CCGGCTGAAG	GCTGAGTGTC	CGGCTGAGGC	CTGAGCGAGT	GTCCAGCCAA
3651	GGGCTGAGTG	TCCAGCACAC	CTGCCGTCTT	CACTTCCCCA	CAGGCTGGCG
3701	CTCGGCTCCA	CCCCAGGGCC	AGCTTTTCCT	CACCAGGAGC	CCGGCTTCCA
3751	CTCCCCACAT	AGGAATAGTC	CATCCCCAGA	TTCGCCATTG	TTCACCCCTC
3801	GCCCTGCCCT	CCTTTGCCTT	CCACCCCCAC	CATCCAGGTG	GAGACCCTGA
3851	GAAGGACCCT	GGGAGCTCTG	GGAATTTGGA	GTGACCAAAG	GTGTGCCCTG
3901	TACACAGGCG	AGGACCCTGC	ACCTGGATGG	GGGTCCCTGT	GGGTCAAATT
3951	GGGGGGAGGT	GCTGTGGGAG	TAAAATACTG	AATATATGAG	TTTTTCAGTT
4001	TTGAAAAAAA	AAAAAAAAA	AAAAAAAA		

FIG. 71B

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		GCAGCGCTGCGTCCTGCTGCGCACGTGGGAAGCCCTGGCCCCGGGCCACCCCCGCGATGCC	CO
	1	CGTCGCGACGCAGGACGACGCGTGCACCCTTCGGGACCGGGGCCGGTGGGGGCGCTACGG	60
a b c		A A L R P A A H V G S P G P G H P R D A Q R C V L L R T W E A L A P A T P A M P S A A S C C A R G K P W P R P P P R C R	<u>-</u> -
	61	GCGCGCTCCCCGCTGCCGAGCCGTGCGCTCCCTGCTGCGCAGCCACTACCGCGAGGTGCT+ CGCGCGAGGGGCGACGCTCGGCACGCGACGCGACGCGTCGGTGATGCCGCTCCACGA	120
a b c		A R S P L P S R A L P A A Q P L P R G A R A P R C R A V R S L L R S H Y R E V L A L P A A E P C A P C C A A T T A R C C	- -
	121	GCCGCTGGCCACGTTCGTGCGGCGCCTGGGGCCCCAGGGCTGGCGGCTGGTGCAGCGCGG + CGGCGACCGGTGCAAGCACGCCGCGGACCCCGGGGTCCCGACCGCCGACCACGTCGCGCC	180
a b c		A A G H V R A A P G A P G L A A G A A R P L A T F V R R L G P Q G W R L V Q R G R W P R S C G A W G P R A G G W C S A G	- - -
	181	GGACCCGGCGCTTTCCGCGCGNTGGTGGCCCANTGCNTGGTGTGCGTGCCCTGGGANGN+ CCTGGGCCGCCGAAAGGCGCGCNACCACCGGGTNACGNACCACACGCACGGGACCCTNCN	240
a b c		G P G G F P R ? G G P ? ? G V R A L G ? D P A A F R A ? V A ? C ? V C V P W ? ? T R R L S A R W W P ? A W C A C P G ? ?	- - -
	241	ANGGCNGCCCCCGCCGCCCCCTCCTTCCGCCAGGTGTCCTGCCTGAANGANCTGGTGGC	300
a b c	241		- -
b		TNCCGNCGGGGGGGGGGGGGAGGAAGGCGGTCCACAGGACGGACTTNCTNGACCACCG	- - -
b		TNCCGNCGGGGGGGGGGGGGAGGAGGCGGTCCACAGGACGACTTNCTNGACCACCG ? A A P R R P L L P P G V L P E ? ? G G ? ? P P A A P S F R Q V S C L ? ? L V A G ? P P P P P S A R C P A * ? ? W W P CCGAGTGCTGCANANGCTGTGCGANCGCGGCGCGAANAACGTGCTGGCCTTCGC	- - - 360
b c a b	301	TNCCGNCGGGGGGGGGGGGGGAGGAGGCGGTCCACAGGACGACTTNCTNGACCACCG ? A A P R R P L L P P G V L P E ? ? G G ? ? P P A A P S F R Q V S C L ? ? L V A G ? P P P P P P S A R C P A * ? ? W W P CCGAGTGCTGCANANGCTGTGCGANCGCGGCGCGAANAACGTGCTGGCCTTCGGCTTCGC GGCTCACGACGTNTNCGACACGCTNGCGCCGCGCTTNTTGCACGACCGGAAGCCGAAGCG P S A A ? A V R ? R R E ? R A G L R L R R V L ? ? L C ? R G A ? N V L A F G F A	- - - 360 - -
b c a b	301	TNCCGNCGGGGGGGGGGGGGGAGGAGGCGGTCCACAGGACGACTTNCTNGACCACCG ? A A P R R P L L P P G V L P E ? ? G G ? ? P P A A P S F R Q V S C L ? ? L V A G ? P P P P P P S A R C P A * ? ? W W P CCGAGTGCTGCANANGCTGTGCGANCGCGGCGCGCAANAACGTGCTGGCCTTCGCCTTCGC GGCTCACGACGTNTNCGACACGCTNGCGCCGCGCTTNTTGCACGACCGGAAGCCGAAGCG P S A A ? A V R ? R R E ? R A G L R L R R V L ? ? L C ? R G A ? N V L A F G F A E C C ? ? C A ? A A R ? T C W P S A S R GCTGCTGGACGGGGCCCCCCCGAGGCCTTCACCACCACCAGCGTGCGAGCTA	- - 360 - - - 420
b c a b c	301 361	TNCCGNCGGGGGGGGGGGGGGGAGAGGCGGTCCACAGGACGACTTNCTNGACCACCG ? A A P R R P L L P P G V L P E ? ? G G ? ? P P A A P S F R Q V S C L ? ? L V A G ? P P P P P P S A R C P A * ? ? W W P CCGAGTGCTGCANANGCTGTGCGANCGCGGCGCGCAANAACGTGCTGGCCTTCGGCTTCGC GGCTCACGACGTNTNCGACACGCTNGCGCCGCGCTTNTTGCACGACCGGAAGCCGAAGCG P S A A ? A V R ? R R E ? R A G L R L R R V L ? ? L C ? R G A ? N V L A F G F A E C C ? ? C A ? A A R ? T C W P S A S R GCTGCTGGACGGGGGCCCCCCGGGGGGCCTTCACCACCACCAGCGTGCGCAGCTA CGACGACCTGCCCCGGGGGGCCCCCCCGAGGCCTTCACCACCACCAGCGTGCGCAGCTA CGACGACCTGCCCCGGGGGGCCCCCCCGGGGGGCTCCGGAAGTGGTGGTCGCACGCGTCGAT A A G R G P R G P P R G L H H Q R A Q L L L D G A R G G P P E A F T T S V R S Y	- - 360 - - - 420

FIG. 72A



a b c		PRGRRAGSPAGTLR??CAGRVGDDVLVHLLARCA?FVLVAWATTCWFTCWHAAR?LCWW	
	541	GGNTCCCAGCTGCGCCTACCANGTGTGCGGGCCGCCGCTGTACCAGCTCGGCGCTGCNAC	600
		CCNAGGGTCGACGCGGATGGTNCACACGCCCGGCGGCGACATGGTCGAGCCGCGACGNTG	
a b c		G S Q L R L P ? V R A A A V P A R R C ? ? P S C A Y ? V C G P P L Y Q L G A A T ? P A A P T ? C A G R R C T S S A L ? L	
	601	TCAGGCCCGGCCCCCCCACACGCTANTGGACCCGAANGCGTCTGGGATCCAACGGGCCT+ AGTCCGGGCCGGGGGGGGTGTGCGATNACCTGGGCTTNCGCAGACCCTAGGTTGCCCGGA	660
a b c		S G P A P A T R ? W T R ? R L G S N G P Q A R P P P H A ? G P E ? V W D P T G L R P G P R H T L ? D P ? A S G I Q R A W	-
	661	GGAACCATAGCGTCAGGGAGGCCGGGGTCCCCCTGGGCTGCCAGCCCCGGGTGCGAGGAG+ CCTTGGTATCGCAGTCCCTCCGGCCCCAGGGGGACCCGACGGTCGGGGCCCACGCTCCTC	720
a b c		G T I A S G R P G S P W A A S P G C E E E P * R Q G G R G P P G L P A P G A R R N H S V R E A G V P L G C Q P R V R G G	-
	721	GCGCGGGGGCAGTGCCGAAGTCTGCCGTTGCCCAAGAGGCCCAGGCGTGGCGCTGC	780
a b c		A R G Q C Q P K S A V A Q E A Q A W R C R G G S A S R S L P L P K R P R R G A A A G A V P A E V C R C P R G P G V A L P	-
	781	CCCTGAGCCGGAGCGGACGCCGTTGGGCAGGGGTCCTGGGCCCACCCGGGCAGGACGCC+ GGGACTCGGCCTGCGGGCAACCCGTCCCCAGGACCCGGGTGGGCCCGTCCTGCGG	840
a b c		P * A G A D A R W A G V L G P P G Q D A P E P E R T P V G Q G S W A H P G R T P L S R S G R P L G R G P G P T R A G R L	-
	841	TGGACCGAGTGACCGTGGTTTCTGTGTGTGTCACCTGCCAGACCCGCCGAAGAAGCCAC+ ACCTGGCTCACTGGCACCAAAGACACACACAGTGGACGGTCTGGGCGCTTCTTCGGTG	900
a b c		W T E * P W F L C G V T C Q T R R R S H G P S D R G F C V V S P A R P A E E A T D R V T V V S V W C H L P D P P K K P P	
	901	CTCTTTGGAGGGTGCGCTCTCTGGCACGCCACTCCCACCCA	960
a b c		L F G G C A L W H A P L P P I R G P P A S L E G A L S G T R H S H P S V G R Q H L W R V R S L A R A T P T H P W A A S T	<u>-</u> -
	961	CCACGCGGGCCCCCATCCACATCGCGGCCACCACGTCCTGGGACACGCCTTGTCCCCCG GGTGCGCCCGGGGGGTAGGTGTAGCGCCGGTGGTGCAGGACCCTGTGCGGAACAGGGGGC	1020

FIG. 72B



a b c		PRGPPIHIAATTSWDTPCPP- HAGPPSTSRPPRPGTRLVPR- TRAPHPHRGHHVLGHALSPG-
	1021	GTGTACGCCGAGACCAAGCACTTCCTCTACTCCTCAGGCGACAAGNACACTGCGNCCCTC
a b c		V Y A E T K H F L Y S S G D K ? T A ? L - C T P R P S T S S T P Q A T ? T L R P S - V R R D Q A L P L L R R Q ? H C ? P P -
	1081	CTTCCTACTCAATATATCTGAGGCCCAGCCTGACTGGCGTTCGGGAGGTTCGTGGAGACA+ 1140 GAAGGATGAGTTATATAGACTCCGGGTCGGACTGACCGCAAGCCCTCCAAGCACCTCTGT
a b c		L P T Q Y I * G P A * L A F G R F V E T - F L L N I S E A Q P D W R S G G S W R ? - S Y S I Y L R P S L T G V R E V R G D ? -
	1141	NTCTTTCTGGTTCCAGGCCTTGGATGCCAGGATTCCCCGCAGGTTGCCCCGCCTGCCCCA
a b c		? F L V P G L G C Q D S P Q V A P P A P - S F W F Q A L D A R I P R R L P R L P Q - L S G S R P W M P G F P A G C P A C P S -
	1201	GCGNTACTGGCAAATGCGGCCCCTGTTTCTGGAGCTGCTTGGGAACCACGCGCAGTGCCC
a b c		A ? L A N A A P V S G A A W E P R A V P - R Y W Q M R P L F L E L L G N H A Q C P - ? T G K C G P C F W S C L G T T R S A P -
	1261	CTACGGGGTGTTCCTCAAGACGCACTGCCCGCTGCGAGCTGCGGTCACCCCAGCAGCCGG
a b c		L R G V P Q D A L P A A S C G H P S S R - Y G V F L K T H C P L R A A V T P A A G - T G C S S R R T A R C E L R S P Q Q P V -
	1321	TGTCTGTGCCCGGGAGAAGCCCCAGGGCTCTGTGGCGGCCCCCGAGGAGGAGGAACACAG+
a b c		C
	1381	ACCCCGTCGCCTGGTGCAGCTGCTCCGCCAGCACAGCAGCCCCTGGCAGGTGTACGGCT+ 1440 TGGGGGCAGCGGACCACGTCGACGAGGCGGTCGTGTCGT
a b c		T P V A W C S C S A S T A A P G R C T A - P P S P G A A A P P A Q Q P L A G V R L - P R R L V Q L L R Q H S S P W Q V Y G F -
	1441	TCGTGCGGGCCTGCGCCGGCTGGTGCCCCCAGGCCTCTGGGGCTCCAGGCACAACG+



S C G P A C A G W C P Q A S G A P G T T - R A G L P A P A G A P R P L G L Q A Q R - V R A C L R R L V P P G L W G S R H N E b ${\tt AACGCCGCTTCCTCAGGAACACCAAGAAGTTCATCTCCCTGGGGAAGCATGCCAAGCTCT}$ 1501 -----+ 1560 TTGCGCGAAGGAGTCCTTGTGGTTCTTCAAGTAGAGGGACCCCTTCGTACGGTTCGAGA b R R F L R N T K K F I S L G K H A K L S -C CGCTGCAGGAGCTGACGTGGAAGATGAGCGTGCGGGACTGCGCTTGGCTGCGCAGGAGCC 1561 -----+ 1620 GCGACGTCCTCGACTGCACCTTCTACTCGCACGCCCTGACGCGAACCGACGCGTCCTCGG R C R S * R G R * A C G T A L G C A G A а A A G A D V E D E R A G L R L A A Q E P -L Q E L T W K M S V R D C A W L R R S Ph C ${\tt CAGGGGTTGGCTGTTCCGGCCGCAGAGCACCGTCTGCGTGAGGAGATCCTGGCCAAGT}$ 1621 -----+ 1680 GTCCCCAACCGACACAGGCCGGCGTCTCGTGGCAGACGCACTCCTCTAGGACCGGTTCA Q G L A V F R P Q S T V C V R R S W P S а R G W L C S G R R A P S A * G D P G Q V b G V G C V P A A E H R L R E E I L A K F -C 1681 -----+ 1740 S C T G * * V C T S S S C S G L S F M S -P A L A D E C V R R R A A Q V F L L C H a b L H W L M S V Y V V E L L R S F F Y V T -C CGGAGACCACGTTTCAAAAGAACAGGCTCTTTTTCTACCGGAAGAGTGTCTGGAGCAAGT 1741 -----+ 1800 GCCTCTGGTGCAAAGTTTTCTTGTCCGAGAAAAAGATGGCCTTCTCACAGACCTCGTTCA а b C ${\tt TGCAAAGCATTGGAATCAGACAGCACTTGAAGAGGGTGCAGCTGCGGGAGCTGTCGGAAG}$ 1801 -----++----++ 1860 ACGTTTCGTAACCTTAGTCTGTCGTGAACTTCTCCCACGTCGACGCCCTCGACAGCCTTC C K A L E S D S T * R G C S C G S C R K a AKHWNQTALEEGAAAGAVGSb O S I G I R O H L K R V O L R E L S E A -CAGAGGTCAGGCAGCCAGGCCAGGCCCGCCTGCTGACGTCCAGACTCCGCTTCA 1861 -----+ 1920 GTCTCCAGTCCGTCGTAGCCCTTCGGTCCGGGCGGACGACTGCAGGTCTGAGGCGAAGT Q R S G S I G K P G P P C * R P D S A S а RGQAASGSQARPADVQTPLHb E V R Q H R E A R P A L L T S R L R F I -C TCCCCAAGCCTGACGGCTGCGGCCGATTGTGAACATGGACTACGTCGTGGGAGCCAGAA 1921 -----+ 1980 ${\tt AGGGGTTCGGACTGCCGACGCCGGCTAACACTTGTACCTGATGCAGCACCCTCGGTCTT}$



a b c		S P S L T G C G R L * T W T T S W E P E - P Q A * R A A A D C E H G L R R G S Q N - P K P D G L R P I V N M D Y V V G A R T -
	1981	CGTTCCGCAGAGAAAGAGGGCCGAGCGTCTCACCTCGAGGGTGAAGGCACTGTTCAGCG+ 2040 GCAAGGCGTCTCTTTTCTCCCGGCTCGCAGAGTGGAGCTCCCACTTCCGTGACAAGTCGC
a b c		R S A E K R G P S V S P R G * R H C S A - V P Q R K E G R A S H L E G E G T V Q R - F R R E K R A E R L T S R V K A L F S V -
	2041	TGCTCAACTACGAGCGGGCGCGCGCCCCGGCCTCCTGGGCCCTCTGTGCTGGGCCTGG
a b c		C S T T S G R G A P A S W A P L C W A W - A Q L R A G A A P R P P G R L C A G P G - L N Y E R A R P G L L G A S V L G L D -
	2101	ACGATATCCACAGGGCCTGGCGCACCTTCGTGCTGCGTGTGCGGGCCCAGGACCCGCCGC
a b c		T I S T G P G A P S C C V C G P R T R R - R Y P Q G L A H L R A A C A G P G P A A - D I H R A W R T F V L R V R A Q D P P P -
	2161	CTGAGCTGTACTTTGTCAAGGTGGATGTGACGGCGCGTACGACACCATCCCCCAGGACA
a b c		L S C T L S R W M * R A R T T P S P R T - * A V L C Q G G C D G R V R H H P P G Q - E L Y F V K V D V T G A Y D T I P Q D R -
	2221	GGCTCACGGAGGTCATCGCCAGCATCATCAAACCCCAGAACACGTACTGCGTCGGT
a b c		G S R R S S P A S S N P R T R T A C V G - A H G G H R Q H H Q T P E H V L R A S V - L T E V I A S I I K P Q N T Y C V R R Y -
	2281	ATGCCGTGGTCCAGAAGGCCGCCCATGGGCACGTCCGCAAGGCCTTCAAGAGCCACGTCT+ 2340 TACGGCACCAGGTCTTCCGGCGGGTACCCGTGCAGGCGTTCCGGAAGTTCTCGGTGCAGA
a b c		M P W S R R P P M G T S A R P S R A T S - C R G P E G R P W A R P Q G L Q E P R L - A V V Q K A A H G H V R K A F K S H V S -
	2341	CTACCTTGACAGACCTCCAGCCGTACATGCGACAGTTCGTGGCTCACCTGCAGGANAACA+ 2400 GATGGAACTGTCTGGAGGTCGGCATGTACGCTGTCAAGCACCGAGTGGACGTCCTNTTGT
a b c		L P * Q T S S R T C D S S W L T C R ? T - Y L D R P P A V H A T V R G S P A G ? Q - T L T D L Q P Y M R Q F V A H L Q ? N S -
	2401	GCCCGCTGAGGGATGCCGTCGTCATCGAGCAGAGCTCCTCCCTGAATGAGGCCAGCAGTG+ 2460 CGGGCGACTCCCTACGGCAGCAGTAGCTCGTCTCGAGGAGGGACTTACTCCGGTCGTCAC

FIG. 72E



A R * G M P S S S S R A P P * M R P A V PAEGCRRHRAELLPE*GQQW b V V I E Q S S S L N E A S S G -PLRDA ${\tt GCCTCTTCGACGTCTTCCTACGCTTCATGTGCCACCACGCCGTGCGCATCAGGGGCAAGT}$ 2461 -------+ 2520 CGGAGAAGCTGCAGAAGGATGCGAAGTACACGGTGGTGCGGCACGCGTAGTCCCCGTTCA A S S T S S Y A S C A T T P C A S G A S - P L R R L P T L H V P P R R A H Q G Q V b L F D V F L R F M C H H A V R I R G K S -C CCTACGTCCAGTGCCAGGGGATCCCGCAGGGCTCCATCCTCTCCACGCTGCTCTGCAGCC 2521 -----+ 2580 GGATGCAGGTCACGGTCCCCTAGGGCGTCCCGAGGTAGGAGAGGTGCGACGAGACGTCGG P T S S A R G S R R A P S S P R C S A A - L R P V P G D P A G L H P L H A A L Q P b Y V Q C Q G I P Q G S I L S T L L C S L -С TGTGCTACGGCGACATGGAGAACAAGCTGTTTGCGGGGATTCGGCGGGACGGGCTGCTCC 2581 -----+ 2640 ACACGATGCCGCTGTACCTCTTGTTCGACAAACGCCCCTAAGCCGCCCTGCCCGACGAGG C A T A T W R T S C L R G F G G T G C S а V L R R H G E Q A V C G D S A G R A A P -C Y G D M E N K L F A G I R R D G L L Lb С TGCGTTTGGTGGATGATTTCTTGTTGGTGACACCTCACCTCACCCACGCGAAAACCTTCC 2641 -----+ 2700 ACGCAAACCACTACTAAAGAACAACCACTGTGGAGTGGGTGCGCTTTTTGGAAGG C V W W M I S C W * H L T S P T R K P S - A F G G * F L V G D T S P H P R E N L P - R L V D D F L L V T P H L T H A K T F L b C TCAGGACCTGGTCCGAGGTGTCCCTGAGTATGGCTGCGTGAACTTGCGGAAGACAG 2701 -----+ 2760 AGTCCTGGGACCAGGCTCCACAGGGACTCATACCGACGCACCACTTGAACGCCTTCTGTC S G P W S E V S L S M A A W * T C G R Q - Q D P G P R C P * V W L R G E L A E D S а b R T L V R G V P E Y G C V V N L R K T V -C TGGTGAACTTCCCTGTAGAAGACGAGGCCCTGGGTGGCACGGCTTTTGTTCAGATGCCGG ${\tt ACCACTTGAAGGGACATCTTCTGCTCCGGGACCCACCGTGCCGAAAACAAGTCTACGGCC}$ W * T S L * K T R P W V A R L L F R C R а GELPCRRGPGWHGFCSDAGb FPVEDEALGGTAFVOMPA- $\tt CCCACGGCCTATTCCCCTGGTGCGGCCTGCTGGATACCCGGACCCTGGAGGTGCAGA$ 2821 -----+-----+ 2880 GGGTGCCGGATAAGGGGACCACGCCGGACGACGACCTATGGGCCTGGGACCTCCACGTCT PTAYSPGAACCWIPGPWRCR PRPIPLVRPAAGYPDPGGAEb HGLFPWCGLLLDTRTLEVQS- \sim GCGACTACTCCAGCTATGCCCGGACCTCCATCAGAGCCAGTCTCACCTTCAACCGCGGCT 2881 -----+ 2940 CGCTGATGAGGTCGATACGGGCCTGGAGGTAGTCTCGGTCAGAGTGGAAGTTGGCGCCGA



a b c		A T T P A M P G P P S E P V S P S T A A R L L Q L C P D L H Q S Q S H L Q P R L D Y S S Y A R T S I R A S L T F N R G F	- - -
29	941	${\tt TCAAGGCTGGGAGGAACATGCGTCGCAAACTCTTTGGGGTCTTGCGGCTGAAGTGTCACAAGTGTCACAAGTGTCCGACCCTCCTTGTACGCAGCGTTTGAGAAACCCCAGAACGCCGACTTCACAGTGT}$	3000
a b c		S R L G G T C V A N S L G S C G * S V T Q G W E E H A S Q T L W G L A A E V S Q K A G R N M R R K L F G V L R L K C H S	- - -
30	001	GCCTGTTTCTGGATTTGCAGGTGAACAGCCTCCAGACGTGTGCACCAACATCTACAAGA 	3060
a b c		A C F W I C R * T A S R R C A P T S T R P V S G F A G E Q P P D G V H Q H L Q D L F L D L Q V N S L Q T V C T N I Y K I	
30	061	TCCTCCTGCTGCAGGCGTACAGGTTTCACGCATGTGTGCTGCAGCTCCCATTTCATCAGC	3120
a b c		S S C C R R T G F T H V C C S S H F I S P P A A G V Q V S R M C A A A P I S S A L L L Q A Y R F H A C V L Q L P F H Q Q	<u>-</u> -
3:	121	$\begin{tabular}{lllllllllllllllllllllllllllllllllll$	3180
a b c		K F G R T P H F S C A S S L T R P P S A S L E E P H I F P A R H L * H G L P L L V W K N P T F F L R V I S D T A S L C Y	- - -
3:	181	ACTCCATCCTGAAAGCCAAGAACGCAGGGATGTCGCTGGGGGCCAAGGGCGCCGGCCCGCCC	3240
a b c		T P S * K P R T Q G C R W G P R A P P A L H P E S Q E R R D V A G G Q G R R R P S I L K A K N A G M S L G A K G A A G P	- - -
32	241	CTCTGCCCTCCGAGGCCGTGCAGTGGCTGTCCACCAAGCATTCCTGCTCAAGCTGACTC+ GAGACGGGAGGCTCCGGCACGTCACCGACACGGTGGTTCGTAAGGACGAGTTCGACTGAG	3300
a b c		L C P P R P C S G C A T K H S C S S * L S A L R G R A V A V P P S I P A Q A D S L P S E A V Q W L C H Q A F L L K L T R	
3:	301	GACACCGTGTCACCTACGTGCCACTCCTGGGGTCACTCAGGACAGCCCAGACGCAGCTGA + CTGTGGCACAGTGGATGCACGGTGAGGACCCCAGTGAGTCCTGTCGGGTCTGCGTCGACT	3360
a b c		$\begin{array}{cccccccccccccccccccccccccccccccccccc$	
3:	361	GTCGGAAGCTCCCGGGGACGACGCTGACTGCCCTGGAGGCCGAGCCAACCCGGCACTGC	3420



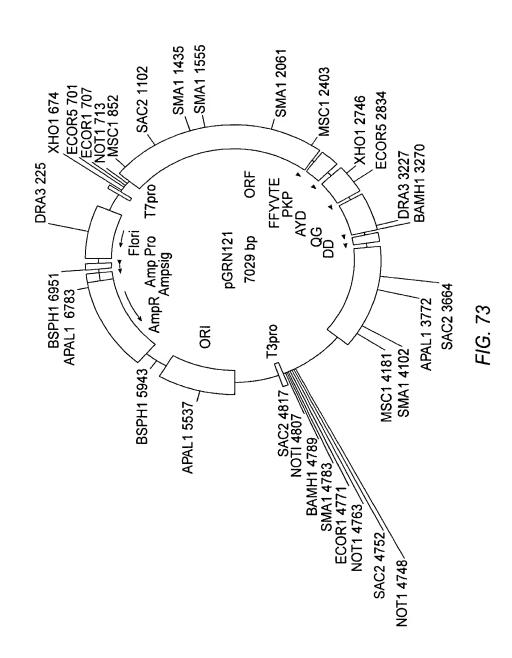
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V G S S R G R R * L P W R P Q P T R H C -
       SEAPGDDADCPGGRSQPGTA-
RKLPGTTLTALEAAANPALP-
b
      \verb|CCTCAGACTTCAAGACCATCCTGGACTGATGGCCACCCGCCCACAGCCAGGCCGAGAGCA|\\
  3421 ------ 3480
      GGAGTCTGAAGTTCTGGTAGGACCTGACTACCGGTGGGCGGGTGTCGGTCCGGCTCTCGT
      P Q T S R P S W T D G H P P T A R P R A L R L Q D H P G L M A T R P Q P G R E Q
b
       S D F K T I L D * W P P A H S Q A E S R -
C
      3481 -----+ 3540
      b
C
      3541 -----+ 3600
      GGTCCGGGCGTGGCGACCCTCAGACTCCGGACTCACTACAAACCGGCTCCGGACGTACA
      P G P H R W E S E A * V S V W P R P A C - Q A R T A G S L R P E * V F G R G L H V - R P A P L G V * G L S E C L A E A C M S -
b
C
      CCGGCTGAAGGCTGACTGTCCGGCTGAGGCCTGAGCGAGTGTCCAGCCAAGGGCTGAGTG
  3601 ------ 3660
      GGCCGACTTCCGACTCACAGGCCGACTCCGGACTCGCTCACAGGTCGGTTCCCGACTCAC
      P A E G * V S G * G L S E C P A K G * V - R L K A E C P A E A * A S V Q P R A E C - G * R L S V R L R P E R V S S Q G L S V -
b
C
      TCCAGCACACCTGCCGTCTTCACTTCCCCACAGGCTGGCGCTCGGCTCCACCCCAGGGCC
  3661 -----+ 3720
      AGGTCGTGTGGACGGCAGAAGTGAAGGGGTGTCCGACCGCGAGCCGAGGTGGGGTCCCGG
      S S T P A V F T S P Q A G A R L H P R A - P A H L P S S L P H R L A L G S T P G P -
а
b
        Q H T C R L H F P T G W R S A P P Q G Q -
      AGCTTTTCCTCACCAGGAGCCCGGCTTCCACTCCCCACATAGGAATAGTCCATCCCCAGA
      {\tt TCGAAAAGGAGTGGTCCTCGGGCCGAAGGTGAGGGGTGTATCCTTATCAGGTAGGGGTCT}
      b
         FLTRSPASTPHIGIVHPQI-
      TTCGCCATTGTTCACCCCTCGCCCTGCCCTCTTTGCCTTCCACCCCCACCATCCAGGTG
  3781 ------ 3840
      AAGCGGTAACAAGTGGGGAGCGGGACGGAGGAAACGGAAGGTGGGGGTGGTAGGTCCAC
      FAIVHPSPCPPLPSTPTIQV
       S P L F T P R P A L L C L P P P P S R W - R H C S P L A L P S F A F H P H H P G G -
b
C
      GAGACCCTGAGAAGGACCCTGGGAGCTCTGGGAATTTGGAGTGACCAAAGGTGTGCCCTG
  CTCTGGGACTCTTCCTGGGACCCTCGAGACCCTTAAACCTCACTGGTTTCCACACGGGAC
```



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E T L R R T L G A L G I W S D Q R C A L - R P * E G P W E L W E F G V T K G V P C - D P E K D P G S S G N L E * P K V C P V -
b
C
        TACACAGGCGAGGACCCTGCACCTGGATGGGGTCCCTGTGGGTCAAATTGGGGGGAGGT
   ATGTGTCCGCTCCTGGGACGTGGACCTACCCCCAGGGACACCCAGTTTAACCCCCCTCCA
        Y T G E D P A P G W G S L W V K L G G G - T Q A R T L H L D G G P C G S N W G E V - H R R G P C T W M G V P V G Q I G G R C -
b
C
        3961 -----+ 4020
        A V G V K Y * I Y E F F S F E K K K K K - L W E * N T E Y M S F S V L K K K K K K - C G S K I L N I * V F Q F * K K K K K K -
b
        AAAAAAAA
   4021 ----- 4029
        \mathbf{TTTTTTTTT}
        K K K
K K
K K
а
b
C
```

FIG. 721







1 met GCAGCGCTGCGTCCTGCGCACGTGGGAAGCCCTGGCCCCGGCCACCCCGCG ATG pro arg ala pro arg cys arg ala val arg ser leu leu arg ser CCG CGC GCT CCC CGC TGC CGA GCC GTG CGC TCC CTG CTG CGC AGC his tyr arg glu val leu pro leu ala thr phe val arg arg leu CAC TAC CGC GAG GTG CTG CCG CTG GCC ACG TTC GTG CGG CGC CTG 40 gly pro gln gly trp arg leu val gln arg gly asp pro ala ala GGG CCC CAG GGC TGG CGG CTG GTG CAG CGC GGG GAC CCG GCG phe arg ala leu val ala gln cys leu val cys val pro trp asp TTC CGC GCG CTG GTG GCC CAG TGC CTG GTG TGC GTG CCC TGG GAC ala arg pro pro pro ala ala pro ser phe arg gln val ser cys GCA CGG CCG CCC CCC GCC CCC TCC TTC CGC CAG GTG TCC TGC leu lys glu leu val ala arg val leu gln arg leu cys glu arg CTG AAG GAG CTG GTG GCC CGA GTG CTG CAG AGG CTG TGC GAG CGC 100 gly ala lys asn val leu ala phe gly phe ala leu leu asp gly GGC GCG AAG AAC GTG CTG GCC TTC GGC TTC GCG CTG CTG GAC GGG 110 ala arg gly gly pro pro glu ala phe thr thr ser val arg ser GCC CGC GGG GGC CCC CCC GAG GCC TTC ACC ACC AGC GTG CGC AGC 130 tyr leu pro asn thr val thr asp ala leu arg gly ser gly ala TAC CTG CCC AAC ACG GTG ACC GAC GCA CTG CGG GGG AGC GGG GCG trp gly leu leu arg arg val gly asp asp val leu val his TGG GGG CTG CTG CGC CGC GTG GGC GAC GAC GTG CTG GTT CAC 160 leu leu ala arg cys ala leu phe val leu val ala pro ser cys CTG CTG GCA CGC TGC GCG CTC TTT GTG CTG GTG GCT CCC AGC TGC ala tyr gln val cys gly pro pro leu tyr gln leu gly ala ala GCC TAC CAG GTG TGC GGG CCG CCG CTG TAC CAG CTC GGC GCT GCC 190 thr gln ala arg pro pro his ala ser gly pro arg arg ACT CAG GCC CGG CCC CCG CCA CAC GCT AGT GGA CCC CGA AGG CGT

FIG. 74A



leu gly cys	200 glu arg GAA CGG	ala trp GCC TGG	asn h	nis ser CAT AGC	val arg GTC AGG	210 glu ala gly GAG GCC GGG
val pro leu GTC CCC CTG	gly leu GGC CTG	pro ala CCA GCC	pro g	220 gly ala GGT GCG	arg arg AGG AGG	arg gly gly CGC GGG GGC
						240 arg arg gly AGG CGT GGC
ala ala pro GCT GCC CCT	glu pro GAG CCG	glu arg GAG CGG	thr p	oro val	gly gln GGG CAG	gly ser trp GGG TCC TGG
ala his pro	260 gly arg GGC AGG	thr arg ACG CGT	gly p	oro ser	asp arg GAC CGT	270 gly phe cys GGT TTC TGT
val val ser GTG GTG TCA	pro ala CCT GCC 290	arg pro AGA CCC	ala g	80 glu glu GAA GAA	ala thr GCC ACC	ser leu glu TCT TTG GAG 300
gly ala leu GGT GCG CTC	ser gly	thr arg ACG CGC	his s CAC T	er his	pro ser CCA TCC	val gly arg GTG GGC CGC
gln his his	ala gly GCG GGC	pro pro	ser t	thr ser	arg pro	pro arg pro CCA CGT CCC
trp asp thr	320 pro cys CCT TGT	pro pro	val t	yr ala AC GCC	glu thr GAG ACC	330 lys his phe AAG CAC TTC
leu tyr ser	ser gly TCA GGC	asp lys GAC AAG	glu g	340 gln leu CAG CTG	arg pro	ser phe leu TCC TTC CTA
leu ser ser CTC AGC TCT	350 leu arg CTG AGG	pro ser CCC AGC	leu t	hr gly	ala arg GCT CGG	360 arg leu val AGG CTC GTG
glu thr ile	phe leu TTT CTG	gly ser GGT TCC	arg p	770 Pro trp	met pro	gly thr pro GGG ACT CCC
arg arg leu CGC AGG TTG	380 pro arg CCC CGC	leu pro	gln a	rg tyr	trp gln TGG CAA	390 met arg pro ATG CGG CCC
leu phe leu CTG TTT CTG	glu leu GAG CTG	leu gly CTT GGG	asn h	00 is ala AC GCG	gln cys CAG TGC	pro tyr gly CCC TAC GGG
val leu leu GTG CTC CTC	410 lys thr AAG ACG	his cys	pro 1 CCG C	eu arg	ala ala GCT GCG	420 val thr pro GTC ACC CCA

FIG. 74B



430 ala ala gly val cys ala arg glu lys pro gln gly ser val ala GCA GCC GGT GTC TGT GCC CGG GAG AAG CCC CAG GGC TCT GTG GCG 440 ala pro glu glu glu asp thr asp pro arg arg leu val gln leu GCC CCC GAG GAG GAG GAC ACA GAC CCC CGT CGC CTG GTG CAG CTG 460 leu arg gln his ser ser pro trp gln val tyr gly phe val arg CTC CGC CAG CAC AGC CCC TGG CAG GTG TAC GGC TTC GTG CGG ala cys leu arg arg leu val pro pro gly leu trp gly ser arg GCC TGC CTG CGC CGG CTG GTG CCC CCA GGC CTC TGG GGC TCC AGG his asn glu arg arg phe leu arg asn thr lys lys phe ile ser CAC AAC GAA CGC CGC TTC CTC AGG AAC ACC AAG AAG TTC ATC TCC leu gly lys his ala lys leu ser leu gln glu leu thr trp lys CTG GGG AAG CAT GCC AAG CTC TCG CTG CAG GAG CTG ACG TGG AAG met ser val arg asp cys ala trp leu arg arg ser pro gly val ATG AGC GTG CGG GAC TGC GCT TGG CTG CGC AGG AGC CCA GGG GTT 530 gly cys val pro ala ala glu his arg leu arg glu glu ile leu GGC TGT GTT CCG GCC GCA GAG CAC CGT CTG CGT GAG GAG ATC CTG 550 ala lys phe leu his trp leu met ser val tyr val val glu leu GCC AAG TTC CTG CAC TGG CTG ATG AGT GTG TAC GTC GTC GAG CTG 560 leu arg ser phe phe tyr val thr glu thr thr phe gln lys asn CTC AGG TCT TTC TTT TAT GTC ACG GAG ACC ACG TTT CAA AAG AAC 580 arg leu phe phe tyr arg pro ser val trp ser lys leu gln ser AGG CTC TTT TTC TAC CGG CCG AGT GTC TGG AGC AAG TTG CAA AGC 590 ile gly ile arg gln his leu lys arg val gln leu arg glu leu ATT GGA ATC AGA CAG CAC TTG AAG AGG GTG CAG CTG CGG GAG CTG 610 ser glu ala glu val arg gln his arg glu ala arg pro ala leu TCG GAA GCA GAG GTC AGG CAG CAT CGG GAA GCC AGG CCC GCC CTG 620 630 leu thr ser arg leu arg phe ile pro lys pro asp gly leu arg CTG ACG TCC AGA CTC CGC TTC ATC CCC AAG CCT GAC GGG CTG CGG 640 pro ile val asn met asp tyr val val gly ala arg thr phe arg CCG ATT GTG AAC ATG GAC TAC GTC GTG GGA GCC AGA ACG TTC CGC

FIG. 74C



650 arg glu lys arg ala glu arg leu thr ser arg val lys ala leu AGA GAA AAG AGG GCC GAG CGT CTC ACC TCG AGG GTG AAG GCA CTG 670 phe ser val leu asn tyr glu arg ala arg arg pro gly leu leu TTC AGC GTG CTC AAC TAC GAG CGG GCG CGC CGC CCC GGC CTC CTG gly ala ser val leu gly leu asp asp ile his arg ala trp arg GGC GCC TCT GTG CTG GGC CTG GAC GAT ATC CAC AGG GCC TGG CGC thr phe val leu arg val arg ala gln asp pro pro pro glu leu ACC TTC GTG CTG CGT GTG CGG GCC CAG GAC CCG CCT GAG CTG 710 tyr phe val lys val asp val thr gly ala tyr asp thr ile pro TAC TTT GTC AAG GTG GAT GTG ACG GGC GCG TAC GAC ACC ATC CCC 730 gln asp arg leu thr glu val ile ala ser ile ile lys pro gln CAG GAC AGG CTC ACG GAG GTC ATC GCC AGC ATC AAA CCC CAG 740 750 asn thr tyr cys val arg arg tyr ala val val gln lys ala ala AAC ACG TAC TGC GTG CGT CGG TAT GCC GTG GTC CAG AAG GCC GCC 760 his gly his val arg lys ala phe lys ser his val ser thr leu CAT GGG CAC GTC CGC AAG GCC TTC AAG AGC CAC GTC TCT ACC TTG 770 thr asp leu gln pro tyr met arg gln phe val ala his leu gln ACA GAC CTC CAG CCG TAC ATG CGA CAG TTC GTG GCT CAC CTG CAG 790 glu thr ser pro leu arg asp ala val val ile glu gln ser ser GAG ACC AGC CCG CTG AGG GAT GCC GTC GTC ATC GAG CAG AGC TCC 800 ser leu asn glu ala ser ser gly leu phe asp val phe leu arg TCC CTG AAT GAG GCC AGC AGT GGC CTC TTC GAC GTC TTC CTA CGC phe met cys his his ala val arg ile arg gly lys ser tyr val TTC ATG TGC CAC CAC GCC GTG CGC ATC AGG GGC AAG TCC TAC GTC 830 gln cys gln gly ile pro gln gly ser ile leu ser thr leu leu CAG TGC CAG GGG ATC CCG CAG GGC TCC ATC CTC TCC ACG CTG CTC 850 cys ser leu cys tyr gly asp met glu asn lys leu phe ala gly TGC AGC CTG TGC TAC GGC GAC ATG GAG AAC AAG CTG TTT GCG GGG 860 ile arg arg asp gly leu leu leu arg leu val asp asp phe leu ATT CGG CGG GAC GGG CTG CTC CTG CGT TTG GTG GAT GAT TTC TTG

FIG. 74D



880 leu val thr pro his leu thr his ala lys thr phe leu arg thr TTG GTG ACA CCT CAC CTC ACC CAC GCG AAA ACC TTC CTC AGG ACC 890 leu val arg gly val pro glu tyr gly cys val val asn leu arg CTG GTC CGA GGT GTC CCT GAG TAT GGC TGC GTG GTG AAC TTG CGG 910 lys thr val val asn phe pro val glu asp glu ala leu gly gly AÁG ACA GTG GTG AAC TTC CCT GTA GAA GAC GAG GCC CTG GGT GGC 920 thr ala phe val gln met pro ala his gly leu phe pro trp cys ACG GCT TTT GTT CAG ATG CCG GCC CAC GGC CTA TTC CCC TGG TGC 940 gly leu leu asp thr arg thr leu glu val gln ser asp tyr GGC CTG CTG CTG GAT ACC CGG ACC CTG GAG GTG CAG AGC GAC TAC ser ser tyr ala arg thr ser ile arg ala ser val thr phe asn TCC AGC TAT GCC CGG ACC TCC ATC AGA GCC AGT GTC ACC TTC AAC 970 arg gly phe lys ala gly arg asn met arg arg lys leu phe gly CGC GGC TTC AAG GCT GGG AGG AAC ATG CGT CGC AAA CTC TTT GGG val leu arg leu lys cys his ser leu phe leu asp leu gln val GTC TTG CGG CTG AAG TGT CAC AGC CTG TTT CTG GAT TTG CAG GTG 1000 asn ser leu gln thr val cys thr asn ile tyr lys ile leu leu AAC AGC CTC CAG ACG GTG TGC ACC AAC ATC TAC AAG ATC CTC CTG leu gln ala tyr arg phe his ala cys val leu gln leu pro phe CTG CAG GCG TAC AGG TTT CAC GCA TGT GTG CTG CAG CTC CCA TTT 1030 his gln gln val trp lys asn pro thr phe phe leu arg val ile CAT CAG CAA GTT TGG AAG AAC CCC ACA TTT TTC CTG CGC GTC ATC 1040 ser asp thr ala ser leu cys tyr ser ile leu lys ala lys asn TCT GAC ACG GCC TCC CTC TGC TAC TCC ATC CTG AAA GCC AAG AAC 1060 ala gly met ser leu gly ala lys gly ala ala gly pro leu pro GCA GGG ATG TCG CTG GGG GCC AAG GGC GCC GCC CCT CTG CCC 1070 ser glu ala val gln trp leu cys his gln ala phe leu leu lys TCC GAG GCC GTG CAG TGG CTG TGC CAC CAA GCA TTC CTG CTC AAG 1090 leu thr arg his arg val thr tyr val pro leu leu gly ser leu CTG ACT CGA CAC CGT GTC ACC TAC GTG CCA CTC CTG GGG TCA CTC

FIG. 74E

1100

AAAAAAAA



arg thr ala gln thr gln leu ser arg lys leu pro gly thr thr AGG ACA GCC CAG ACG CAG CTG AGT CGG AAG CTC CCG GGG ACG ACG 1120 leu thr ala leu glu ala ala ala asn pro ala leu pro ser asp CTG ACT GCC CTG GAG GCC GCA GCC AAC CCG GCA CTG CCC TCA GAC 1130 1132 phe lys thr ile leu asp OP TTC AAG ACC ATC CTG GAC TGA TGGCCACCCGCCCACAGCCAGGCCGAGAGCAGA AGGCCGCACCGCTGGGAGTCTGAGGCCTGAGTGTTTTGGCCGAGGCCTGCATGTCC GGCTGAAGGCTGAGTGTCCGGCTGAGGCCTGAGCGAGTGTCCAGCCAAGGGCTGAGTGTC CAGCACACCTGCCGTCTTCACTTCCCCACAGGCTGGCGCTCCGCCTCCACCCCAGGGCCAG CTTTTCYTCACCAGGAGCCCGGCTTCCACTCCCCACATAGGAATAGTCCATCCCCAGATT CGCCATTGTTCACCCYTCGCCCTGCCYTCCTTTGCCTTCCACCCCCACCATCCAGGTGGA GACCCTGAGAAGGACCCTGGGAGCTCTGGGAATTTGGAGTGACCAAAGGTGTGCCCTGTA CACAGGCGAGGACCCTGCACCTGGATGGGGGTCCCTGTGGGTCAAATTGGGGGGAGGTGC

FIG. 74F

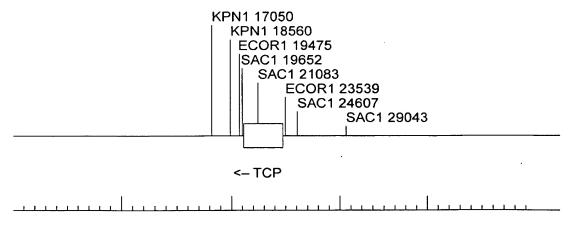


FIG. 75